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INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification 6: C07K 5/11, 14/81, A61K 38/07, 38/57

(11) International Publication Number:

WO 99/51624

(43) International Publication Date:

14 October 1999 (14.10.99)

(21) International Application Number:

PCT/US99/07776

(22) International Filing Date:

8 April 1999 (08.04.99)

(30) Priority Data:

60/081,034

8 April 1998 (08.04.98)

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Published

With international search report.

Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(54) Title: REAGENTS AND METHODS FOR INHIBITING FURIN PROTEASE ACTIVITY

This invention relates to methods and reagents for inhibiting furin endoprotease activity and specifically for inhibiting furin endoprotease-mediated maturation of bioactive proteins in vivo and in vitro. The invention specifically provides peptides, peptide analogues, peptide derivatives and peptido-, organo- and chemical mimetics of said peptide inhibitors of furin endoprotease activity. Methods for using furin endoprotease inhibition to attenuate or prevent viral protein maturation, and thereby alleviate viral infections, are provided. Also provided are methods for using furin endoprotease inhibition to attenuate or prevent proteolytic processing of bacterial toxins, thereby alleviating bacterial infections methods are also provided to inhibit proteolytic processing biologically active proteins and peptides. The invention also provides pharmaceutically acceptable compositions of therapeutically effective amounts of furin endoprotease inhibitors.

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REAGENTS AND METHODS FOR INHIBITING FURIN PROTEASE ACTIVITY

This application claims priority to U.S. Serial No. 60/081,034, filed April 8, 1998.

This invention was made with government support under DK44629 and DK37274 from the National Institutes of Health. The government has certain rights in the invention.

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BACKGROUND OF THE INVENTION

1. Field of the Invention

This invention relates to endoproteases, particularly a novel endoprotease termed furin endoprotease. The invention specifically relates to inhibitors of furin endoprotease activity. In particular, the invention relates to peptides and peptide mimetics derived from a novel variant of 1-antitrypsin that specifically inhibit furin endoprotease activity. The invention also provides methods for using such inhibitors to attenuate or prevent biological proteolytic maturation of bioactive proteins and peptides in vivo and in vitro, in particular with respect to viral proteins and bacterial toxins. Therapeutic methods and pharmaceutical compositions of such inhibitors are also provided directed towards the alleviation and treatment of disease having microbiological etiology.

2. Background of the Related Art

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Most biologically active peptides and proteins are synthesized initially as larger, inactive precursor proteins that are endoproteolytically cleaved during transit through the secretory pathway in the Golgi apparatus in cells expressing such proteins (see Barr, 1991, Cell 66: 1-3 for review). This system comprises an important common mechanism required for synthesis of biologically active proteins and peptides in yeast (Fuller et al., 1988, Ann. Rev. Physiol. 50: 345-362), invertebrates (Scheller et al., 1983, Cell 32: 7-22) and mammalian cells (Sossin et al., 1989, Neuron 2: 1407-1417). Examples of peptides and proteins produced in vivo by exocytotic transport through the

Golgi are precursors of peptide hormones, neuropeptides, growth factors, coagulation factors, serum albumin, cell surface receptors, and adhesion molecules.

Morrison et al., 1985, J. Virol. <u>53</u>: 851-857 disclose that F protein of Newcastle disease virus is processed through the exocytotic transport pathway in infected cells.

Perez & Hunter, 1987, J. Virol. 61: 1069-1614 disclose that the Rous sarcoma virus (RSV) glycoprotein is processed through the exocytotic transport pathway in infected cells.

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Yamada et al., 1988, Virology 165: 268-273 disclose that F protein of mumps virus is processed through the exocytotic transport pathway in infected cells.

Randolph et al., 1990, Virology 174: 450-458 disclose that the prM protein of flaviviruses is processed through the exocytotic transport pathway in infected cells.

A common structural feature of molecules processed through the exocytotic transport pathway is the presence of basic residues or pairs of basic residues at the proteolytic processing site in the molecule. Examples include serum factors (Factors IX; Bentley et al., 1986, Cell 45:343-348; proalbumin; Knowlese et al., 1980, Science 209: 497-499; pro-von Willibrand factor; Bonthron et al., 1986, Nature 324: 270-273), viral polyproteins (human immunodeficiency virus (HIV) gp160; McCune et al., 1988, Cell 53: 55-67; RSV envelope protein; Perez & Hunter, 1987, J. Virol. 61: 1609-1614; yellow fever virus protein; Rice et al., 1985, Science 229: 726-733; measles virus protein; Richardson et al., 1986, Virology 155: 508-523; mumps virus protein; Waxham et al., 1987, Virology 159: 381-389; human cytomegalovirus protein; Spaete et al., 1990, J. Virol. 64: 2922-2931; varicella zooster virus protein; Keller et al., 1986, Virology 152: 181-191), growth factors (preprotransforming growth factor β; Gentry et al., 1988, Molec. Cell. Biol. 8:4162-4168; epidermal growth factor; Gray et al., 1983, Nature 303: 722-725; pro-β-nerve growth factor (NGF); Edwards et al., 1988, Molec. Cell Biol. 8: 2456-2464), receptors (insulin receptor; Yoshimasa et al., 1988, Science 240: 784-787); and bacterial toxins (see Stephen & Pietrowski, 1986, Bacterial Toxins, 2d ed. (Amer. Soc. Microbiol. Washington, D.C.) for review; anthrax toxin; Singh et al., 1989, J. Biol. Chem. 264: 11099-11102). The proteolytic processing site has been identified in some of these molecules.

Berger & Shooter, 1977, *Proc. Natl. Acad. Sci. USA* 74: 3647-3651 disclose the sequence –RSKR- at the proteolytic processing site of pro-β-NGF.

Bentley et al., 1986, ibid., disclose the sequence –RPKR- at the proteolytic processing site of the blood coagulation factor protein Factor IX.

McCune *et al.*, 1988, *ibid.*, disclose the sequence –REKR- at the proteolytic processing site of HIV gp160.

Clepak et al., 1988, Biochem. Biophys. Res. Comm. 157: 747-754 disclose the sequence -RVRR- at the proteolytic processing site of diphtheria toxin.

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Vey et al., 1992, Virology 188: 408-413 disclose the sequence –RX(R/K)R- at the proteolytic processing site of influenza hemagglutinin.

Ogata et al., 1990, J. Biol. Chem. 265: 20678-20685 disclose the sequence -RSKR- at the proteolytic processing site of Pseudomonas exotoxin A.

Klimpel et al., 1992, Proc. Natl. Acad. Sci. USA 89: 10277-10281 disclose the sequence -RX(R/K)R- at the proteolytic processing site of anthrax protective antigen.

A cellular endoprotease termed furin has been identified that specifically recognizes the recognition sequence of proteins processed through the exocytotic secretory pathway (Wise et al., 1990, Proc. Natl. Acad. Sci. USA 87: 9378-9382; Bresnanhan et al., 1990, J. Cell Biol. 111: 2851-2859). This endoprotease is a subtilisin-related, calcium-dependent serine protease (Bresnahan et al., ibid.). A complementary DNA copy of the mRNA encoding this endoprotease has been isolated (Wise et al., ibid.) and sequenced (van den Ouweland et al., 1992, Nucleic Acids Res. 18: 664) and expressed in heterologous cells (Bresnahan et al., ibid). These studies have localized furin by fluoresence immunohistochemistry to the Golgi apparatus of cells expressing this endoprotease (Bresnahan et al., ibid.). Furin has been shown to be capable of proteolytically cleaving a number of exocytotically processed proteins.

Bresnahan et al., ibid., disclosure furin-mediated cleavage of pro-β-NGF.

Wise et al., ibid., disclose furin-mediated cleavage of pro-von Willibrand factor and complement factor C3.

Hosaka et al., 1991, J. Biol, Chem. 266: 12127-12130 disclose furin-mediated cleavage of renin.

Steineke-Grober et al., 1992, EMBO J. 11: 2407-2414 disclose furin-mediated cleavage of influenza hemagglutinin.

Klimpel et al., 1992, Proc. Natl. Acad. Sci. USA 89: 10277-10281 disclose furinmediated cleavage of anthrax protective antigen.

Molloy et al., 1992, J. Biol. Chem. 267: 16396-16402 disclose furin-mediated cleavage of anthrax protective antigen.

Klimpel et al., 1992, Annual Meeting, Amer. Soc. Microbiol. Abst. B-32 disclose furin-mediated cleavage of diphtheria toxin.

One of the present inventors has discovered a mutated variant of α_1 -antitrypsin that effectively inhibits furin endoprotease, termed α_1 -antitrypsin Portland (also termed PDX; SEQ ID No.: 1), as disclosed in U.S. Patent 5,604,210, issued February 18, 1997 and International Application, Publication No. WO 94/16073, published July 21, 1994, the complete disclosure of each of which are explicitly incorporated herein in its entirety. This variant has been genetically-engineered to contain Ala₃₅₅ -> Arg₃₅₅ and Met₃₅₈ -> Arg₃₅₈ mutations, whereby the native sequence of α_1 -antitrypsin is changed from -Ala₃₅₅-Ile-Pro-Met₃₅₈- (SEQ ID No. 2) to -Arg₃₅₅-Ile-Pro-Arg₃₅₈- (SEQ ID No. 3) in the Portland variant.

Furin can also be inhibited by specific peptidyl chloroalkylketones (Garten et al., 1989, Virology 172: 25-31; Molloy et al., ibid.; Hallenberger et al., 1992, Nature 360: 358-361), but these substances are toxic in vivo.

In view of the importance of furin endoprotease in activation of bacterial toxins, viral structural proteins and bioactive molecules, there is a need for the development of safe and specific furin inhibitors for prophylaxis, therapy and biological regulation.

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SUMMARY OF THE INVENTION

This invention provides safe, specific and effective inhibitors of furin endoprotease that are peptides and peptide mimetics of novel variants of the naturally-occurring protease inhibitor, a₁-antitrypsin (Heeb et al., 1990, J. Biol. Chem. 265: 2365-2369; Schapira et al., 1987, J. Clin. Invest. 80:582-585). The peptides and peptide mimetics comprise the sequence Arg-Xaa-Xaa-Arg (SEQ ID No. 4) or peptido-, organo-or chemical mimetics thereof.

The invention provides methods for inhibiting bacterial infection of human cells comprising contacting such cells with an effective amount of a peptide or peptide mimetic of the invention. In a preferred embodiment, the bacterial infection is caused by *Corynebacterium diptheriae*. In another preferred embodiment, the bacterial

infection is caused by *Bacillus anthracis*. In yet another preferred embodiment, the bacterial infection is caused by *Pseudomonas aeruginosa*.

The invention also provides a method of inhibiting bacterial infection in a human comprising administering a therapeutically effective amount of a peptide or peptide mimetic of the invention in a pharmaceutically acceptable carrier. In a preferred embodiment, the bacterial infection is caused by *Corynebacterium diptheriae*. In another preferred embodiment, that bacterial is caused by *Bacillus anthracis*. In yet another preferred embodiment, the bacterial infection is caused by *Pseudomonas aeruginosa*.

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Another method provided by the invention for treating humans with a bacterial infection comprises administering a combination of a therapeutically effective amount of a peptide or peptide mimetic of the invention and a therapeutically effective amount of a second antibacterial compound in a pharmaceutically acceptable carrier. In a preferred embodiment, the bacterial infection is caused by *Corynebacterium diptheriae*. In another preferred embodiment, the bacterial infection is caused by *Bacillus anthracis*. In yet another preferred embodiment, the bacterial infection is caused by *Pseudomonas aeruginosa*.

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Pharmaceutically acceptable compositions effective according to the methods of the invention, comprising a therapeutically effective amount of a peptide or peptide mimetic of the invention capable of blocking endoproteolytic activation of bacterial toxins and a pharmaceutically acceptable carrier or diluent, are also provided.

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The invention provides a method of inhibiting viral infection of human cells comprising contacting such cells with an effective amount of a peptide or peptide mimetic of the invention. In preferred embodiments, the viral infection is caused by human cytomegalovirus (HCMV), yellow fever virus, measles virus, mumps virus, influenza virus, varicella zooster virus, or human immunodeficiency virus (HIV-1). In another preferred embodiment, the human cells are hematopoietic cells, most preferably T lymphocytes.

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The invention also provides a method for inhibiting viral infection in an animal, most preferably a human, comprising administering a therapeutically effective amount of a peptide or peptide mimetic of the invention in a pharmaceutically acceptable carrier. In preferred embodiments, the viral infection is caused by human cytomegalovirus,

yellow fever virus, measles virus, mumps virus, influenza virus, varicella zooster virus, or human immunodeficiency virus.

The invention provides a method of treating humans infected with a virus comprising administering a therapeutically effective amount of a peptide or peptide mimetic of the invention in a pharmaceutically acceptable carrier. In preferred embodiments, the viral infection is caused by human cytomegalovirus, yellow fever virus, measles virus, mumps virus, influenza virus, varicella zooster virus, or human immunodeficiency virus.

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The invention provides a method of treating humans infected with a virus comprising administering a combination of a therapeutically effective amount of a peptide or peptide mimetic of the invention and a therapeutically effective amount of a second antiviral compound in a pharmaceutically acceptable carrier. In preferred embodiment, the virus is HIV-1 and the second antiviral compound is azidothymidine. In another preferred embodiment, the virus is HCMV and the second antiviral compound is foscarnet, gancyclovir, or cidofovir.

Pharmaceutically acceptable compositions effective according to the methods of the invention, comprising a therapeutically effective amount of a peptide or peptide mimetic of the invention having antiviral properties and a pharmaceutically acceptable carrier or diluent, are also provided.

The invention also provides a method of inhibiting proteolytic processing of a biologically active protein or peptide in a cell comprising contacting such cells with a peptide or peptide mimetic of the invention. Preferred biologically active proteins are pro-β-nerve growth factor, blood coagulation factor protein Factor IX, pro-von Willibrand factor, complement factor C3 and renin.

Specific preferred embodiments of the present invention will become evident from the following more detailed description of certain preferred embodiments and the claims.

BRIEF DESCRIPTION OF THE DRAWINGS

Figure 1A is a computer generated molecular model of the reactive site loop (RSL) portion of the α_1 -antitrypsin Portland (PDX) variant comprising the sequence - Arg_{355} -Ile₃₅₆-Pro₃₅₇-Arg₃₅₈- (SEQ ID No. 3). Figure 1A shows a view in which certain

atoms in the peptide backbone and arginine sidechains have been numbered. Figure 1B shows rotated views from six perspectives of the molecule.

Figure 2 shows the structure of *Pseudomonas aeruginosa* pro-exotoxin A (PEA) and human cytomegalovirus pro-gB (HCMV progB) proteins, including the furin recognition site.

Figure 3 sets forth a schematic diagram of an assay using detection of *P. aeruginosa* exotoxin A-mediated cellular protein synthesis inhibition to detect compounds that inhibit furin-mediated maturation of the exotoxin.

Figure 4 is a graph of the results of the assay described in Example 1 and performed according to the protocol set forth in Figure 3.

Figure 5 is a diagram of the cleavage pattern of HCMV pro-gB protein.

Figure 6 sets forth a schematic diagram of an assay using detection of plaque formation in naive human foreskin fibroblasts (HFF) incubated with the supernatant fluid of an HCMV-infected culture of U373 cells in the presence or absence of a putative furin inhibiting compound.

Figure 7 is a graph of the results of a plaque-forming assay described in Example 1 and performed according to the protocol set forth in Figure 6.

Figure 8 is a Western blot of proteins produced by HCMV-infected U373 cells and probed with a gB-specific antibody.

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DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS

For the purposes of this invention, the terms "mimetic," "peptide mimetic," "peptidomimetic," "organomimetic" and "chemical mimetic" are intended to encompass peptide derivatives, peptide analogues and chemical compounds having an arrangement of atoms is a three-dimensional orientation that is equivalent to that of a peptide having the sequence Arg-Xaa-Xaa-Arg (SEQ ID No.: 4). It will be understood that the phrase "equivalent to" as used herein is intended to encompass compounds having substitution of certain atoms or chemical moieties in said peptide with moieties having bond lengths, bend angles and arrangements thereof in the mimetic compound that produce the same or sufficiently similar arrangement or orientation of said atoms and moieties to have the biological function of inhibiting furin endoprotease activity. In the peptide mimetics of the invention, the three-dimensional arrangement of the chemical constituents is

structurally and/or functionally equivalent to the three-dimensional arrangement of the peptide backbone and component amino acid sidechains in the peptide, resulting in such peptido-, organo- and chemical mimetics of the peptides of this invention having substantial biological activity, specifically furin protease inhibiting activity. These terms are used according to the understanding in the art, as illustrated for example by Fauchere, 1986, Adv. Drug Res. 15: 29; Veber & Freidinger, 1985, TINS p.392; and Evans et al., 1987, J. Med. Chem. 30: 1229, incorporated herein by reference.

It is understood that a pharmacophore exists for the biological activity of the PDX protein of the invention, said pharmacophore being defined by the -Arg-Xaa-Xaa-Arg- (SEQ ID No. 4) portion of the PDX protein at residues 355-358. A pharmacophore is understood in the art as comprising an idealized, three-dimensional definition of the structural requirements for biological activity. The determination of the three-dimensional structure of the PDX protein provides the structural information to enable production of peptido-, organo- and chemical mimetics of the functional portion of PDX. Peptido-, organo- and chemical mimetics can be designed to fit each pharmacophore with current computer modeling software (computer aided drug design), as described in more detail below. Said mimetics are produced by structure-function analysis, based on the positional information from the crystallographic-derived molecular positional information disclosed herein.

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In a first embodiment, the invention provides peptides defined by the sequence Arg-Xaa-Xaa-Arg. In a preferred embodiment, the peptides of the invention have the sequence Arg-Xaa-Pro-Arg (SEQ ID No. 5). In additional preferred embodiments, the peptide comprises the sequence Arg-Ile-Pro-Arg (SEQ ID No. 4). It will be understood that the use of "Xaa" for these residues is intended to indicate that any amino acid residue can be substituted for Ile or Pro in these positions with no change in the critical dimensions of the rigid reactive site loop (RSL) of the PDX protein, as disclosed herein. Peptide analogues of the invention include embodiments whereby either of the arginine residues are substituted by positively-charged amino acids including lysine, homolysine, hydroxylysine, ornithine, citrulline and canavanine. In preferred embodiments, both arginine residues are substituted by the same alternative amino acid.

Alternative embodiments of the peptides of the invention include peptides having the sequence B-(Arg-Xaa-Xaa-Arg)-C, wherein "B" and "C" represent amino acid sequences each independently comprising from about 1 to about 40 amino acids, more

preferably from about 5 to about 30 amino acids and most preferably from about 10 to about 25 amino acid residues. In particularly preferred embodiments, the B-(Arg-Xaa-Xaa-Arg)-C peptides of the invention are conformationally-restricted, for example by cyclization or disulfide bond formation, wherein the term "disulfide bond" is intended to encompass sulfide linkages and other disulfide derivatives, particularly those that are more stable than naturally-occurring disulfide bonds. In additional preferred embodiments, the peptides of the invention are derivatized by attachment of sugar moieties to produce glycosylated analogues thereof, wherein the sugar moieties are covalently linked to an asparagine residue ("N-linked" glycosylation) or serine, hydroxyproline, hydroxylysine, or threonine residues ("O-linked" glycosylation). Peptides wherein either the amino or carboxyl termini are derivitized are also within the scope of the peptides of the invention.

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Conjugation with sugars (preferably glucose, glucosamine, galactose, galactosamine, mannose, mannosamine, maltose and the like) at the – or C-terminus, at internal asparagine, serine or threonine residues, or at the N-terminus by means of a succinyl linker, serves to stabilize certain conformational motifs, and may increase solubility and bioavailability of the peptide. Similarly, polyethylene glycol (PEG) conjugation at – or C-terminus or on side chains can stabilize peptides and improve pharmacological performance (see, for example, Delgado et al., 1992, Crit. Rev. Ther. Drug Carrier Syst. 9: 249-304). Such modifications may change the in vitro profile of a mimetic, and enhance circulating half-life and in vivo activity.

Peptides as provided by the invention may be advantageously synthesized by any of the chemical synthesis techniques known in the art, particularly solid-phase synthesis techniques, for example, using commercially-available automated peptide synthesizers. The mimetics of the present invention can be synthesized by solid phase or solution phase methods conventionally used for the synthesis of peptides (see, for example, Merrifield, 1963, J. Amer. Chem. Soc. 85: 2149-54; Carpino, 1973, Acc. Chem. Res. 6: 191-98; Birr, 1978, Aspects of the Merrifield Peptide Synthesis, Springer-Verlag: Heidelberg; The Peptides: Analysis, Synthesis, Biology, Vols. 1, 2, 3, 5, (Gross & Meinhofer, eds.), Academic Press: New York, 1979; Stewart et al., 1984, Solid Phase Peptide Synthesis, 2d. ed., Pierce Chem. Co.: Rockford, Ill.; Kent, 1988, Ann. Rev. Biochem. 57: 957-89; and Gregg et al., 1990, Int. J. Peptide Protein Res. 55: 161-214, which are incorporated herein by reference in their entirety.)

The use of solid phase methodology is preferred. Briefly, an N-protected Cterminal amino acid residue is linked to an insoluble support such as divinylbenzene cross-linked polystyrene, polyacrylamide resin, Kieselguhr/polyamide (pepsyn K). controlled pore glass, cellulose, polypropylene membranes, acrylic acid-coated polyethylene rods or the like. Cycles of deprotection, neutralization and coupling of successive protected amino acid derivatives are used to link the amino acids from the Cterminus according to the amino acid sequence. For some synthetic peptides, an Fmoc strategy using an acid-sensitive resin may be used. Preferred solid supports in this regard are divinylbenzene cross-linked polystyrene resins, which are commercially available in a variety of functionalized forms, including chloromethyl resin, hydroxymethyl resin, paraacetamidomethyl resin, benzhydrylamine (BHA) resin, 4methylbenzhydrylamine (MBHA) resin, oxime resins, 4-alkoxybenzyl alcohol resin (Wang resin), 4-(2',4'-dimethoxyphenylaminomethyl)-phenoxymethyl resin, 2,4dimethoxybenzhydrylamine resin, and 4-(2',4'-dimethoxyphenyl-FMOC-aminomethyl)phenoxyacetamidonorleucyl-MBHA resin (Rink amide MBHA resin). In addition, acidsensitive resins also provide C-terminal acids, if desired.

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A particularly preferred protecting group for alpha, amino acids is base-labile 9fluorenylmethoxycarbonyl (Fmoc). Suitable protecting groups for the side chain functionalities of amino acids chemically compatible with Boc (t-butyloxycarbonyl) and Fmoc groups are well known in the art. When using Fmoc chemistry, the following protected amino acid derivatives are preferred: Fmoc-Cys(Trit), Fmoc-Ser(But), Fmoc-Asn(Trit), Fmoc-Leu, Fmoc-Thr(Trit), Fmoc-Val, Fmoc-Gly, Fmoc-Lys(Boc), Fmoc-Gln(Trit), Fmoc-Glu(OBut), Fmoc-His(Trit), Fmoc-Tyr(But), Fmoc-Arg(PMC(=2,2,5,7,8-pentamethylchroman-6-sulfonyl)), Fmoc-Arg(Boc), Fmoc-Pro, and Fmoc-Trp(Boc). The amino acid residues can be coupled by using a variety of coupling agents and chemistries known in the art, such as direct coupling with DIC (diisopropyl-carbodiimide), DCC (dicyclohexylcarbodiimide), BOP (benzotriazolyl-Noxytrisdimethylaminophosphonium hexa-fluorophosphate), PyBOP (benzotriazole-1-yloxy-tris-pyrrolidinophosphonium hexafluoro-phosphate), or PyBrOP (bromo-trispyrrolidinophosphonium hexafluorophosphate); via preformed symmetrical anhydrides; via active esters such as pentafluorophenyl esters; via performed HOBt (1hydroxybenzotriazole) active esters; by using Fmoc-amino acid fluoride and chlorides; or by using Fmoc-amino acid-N-carboxy anhydrides. Activation with HBTU (2-(1H-

benzotriazole-1-yl)-1,1,3,3-tetramethyluronium hexafluorophosphate) or HATU (2-(1H-7-aza-benzotriazole-1-yl)-(1,1,3,3-tetramethyluronium hexafluorophosphate) in the presence of HOBt or HOAt (7-azahydroxybenztriazole) is preferred.

Solid phase methods can be carried out manually, although automated synthesis on a commercially available peptide synthesizer (e.g., Applied Biosystems 431A or the like; Applied Biosystems, Foster City, CA) is preferred. In a typical synthesis, the first (C-terminal) amino acid is loaded on the chlorotrityl resin. Successive deprotection (with 20% piperidine/NMP (N-methylpyrrolidone)) and coupling cycles according to ABI FastMoc protocols (ABI user bulletins 32 and 33, Applied Biosystems) are used to build the complete peptide sequence. Double and triple coupling, with capping by acetic anhydride, may also be used.

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The synthetic mimetic peptide is advantageously cleaved from the resin and deprotected by treatment with TFA (trifluoroacetic acid) containing appropriate scavengers. Many such cleavage reagents, such as Reagent K (0.75 g crystalline phenol/ 0.25 mL ethanedithiol/ 0.5 mL thioanisole/ 0.5 mL deionized water/ 10 mL TFA), can be used. The peptide is separated from the resin by filtration and isolated by ether precipitation. Further purification may be achieved by conventional methods, such as gel filtration and reverse phase HPLC (high performance liquid chromatography). Synthetic mimetics according to the present invention may be in the form of pharmaceutically acceptable salts, especially base-addition salts and including salts of organic bases and inorganic bases. The base-addition salts of the acidic amino acid residues are prepared by treatment of the peptide with the appropriate base, according to procedures well known to those skilled in the art, or the desired salt may be obtained directly by lyophilization out of the appropriate base solution.

Generally, those skilled in the art will recognize that peptides as described herein may be modified by a variety of chemical techniques to produce compounds having essentially the same activity as the unmodified peptide, and optionally having other desirable properties. For example, carboxylic acid groups of the peptide may be provided in the form of a salt of a pharmaceutically-acceptable cation. Amino groups within the peptide may be in the form of a pharmaceutically-acceptable acid addition salt, such as the HCl, HBr, acetic, benzoic, toluene sulfonic, maleic, tartaric and other organic salts, or may be converted to an amide. Thiols can be protected with any one of a number of well-recognized protecting groups, such as acetamide groups. Those

skilled in the art will also recognize methods for introducing cyclic structures into the peptides of this invention so that the native binding configuration will be more nearly approximated. For example, a carboxyl terminal or amino terminal cysteine residue can be added to the peptide, so that when oxidized the peptide will contain a disulfide bond, thereby generating a cyclic peptide. Other peptide cyclizing methods include the formation of thioethers and carboxyl- and amino-terminal amides and esters.

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Specifically, a variety of techniques are available for constructing peptide derivatives and analogues with the same or similar desired biological activity as the corresponding peptide compound but with more favorable activity than the peptide with respect to solubility, stability, and susceptibility to hydrolysis and proteolysis. Such derivatives and analogues include peptides modified at the N-terminal amino group, the C-terminal carboxyl group, and/or changing one or more of the amido linkages in the peptide to a non-amido linkage. It will be understood that two or more such modifications can be coupled in one peptide mimetic structure (e.g., modification at the C-terminal carboxyl group and inclusion of a -CH₂- carbamate linkage between two amino acids in the peptide, for example).

Amino terminus modifications include but are not limited to alkylating, acetylating, adding a carbobenzoyl group, and forming a succinimide group. Specifically, the N-terminal amino group can be reacted to form an amide group of the formula RC(O)NH-, where R is alkyl, preferably lower alkyl, and is added by reaction with an acid halide or acid anhydride. Typically, the reaction can be conducted by contacting about equimolar or excess amounts (e.g., about 5 equivalents) of an acid halide to the peptide in an inert diluent (e.g., dichloromethane) preferably containing an excess (e.g., about 10 equivalents) of a tertiary amine, such as diisopropylethylamine, to scavenge the acid generated during reaction. Reaction conditions are otherwise conventional (e.g., room temperature for 30 minutes). Alkylation of the terminal amino to provide for a lower alkyl N-substitution followed by reaction with an acid halide as described above will provide for N-alkyl amide group of the formula RC(O)NR-.

Alternatively, the amino terminus can be covalently linked to succinimide group by reaction with succinic anhydride. An approximately equimolar amount or an excess of succinic anhydride (e.g., about 5 equivalents) are used and the terminal amino group is converted to the succinimide by methods well known in the art including the use of an excess (e.g., ten equivalents) of a tertiary amine such as diisopropylethylamine in a

suitable inert solvent (e.g., dichloromethane), as described in Wollenberg et al., U.S. Pat. No. 4,612,132 which is incorporated herein by reference in its entirety. It will also be understood that the succinic group can be substituted with, for example, C₂-through C₆-(i.e., lower) alkyl or --SR substituents (where R is alkyl, preferably lower alkyl), which are prepared in a conventional manner to provide for substituted succinimide at the Nterminus of the peptide. Such alkyl substituents are prepared by reaction of a lower olefin (C₂- through C₆- alkyl) with maleic anhydride in the manner described by Wollenberg et al., supra., and --SR substituents are prepared by reaction of RSH with maleic anhydride. In other advantageous embodiments, the amino terminus is derivatized to form a benzyloxycarbonyl-NH-- or a substituted benzyloxycarbonyl-NH-group. This derivative is produced by reaction with approximately an equivalent amount or an excess of benzyloxycarbonyl chloride (CBZ-Cl) or a substituted CBZ-Cl in a suitable inert diluent (e.g., dichloromethane) preferably containing a tertiary amine to scavenge the acid generated during the reaction. In yet another derivative, the Nterminus comprises a sulfonamide group by reaction with an equivalent amount or an excess (e.g., 5 equivalents) of R-S(O), Cl in a suitable inert diluent (dichloromethane) to convert the terminal amine into a sulfonamide, where R is alkyl and preferably lower alkyl. Preferably, the inert diluent contains excess tertiary amine (e.g., 10 equivalents) such as diisopropylethylamine, to scavenge the acid generated during reaction. Reaction conditions are otherwise conventional as described above. Carbamate groups are produced at the amino terminus by reaction with an equivalent amount or an excess (e.g., 5 equivalents) of ROC(O)Cl or R-OC(O)OC₆H₄-p-NO₂ in a suitable inert diluent (e.g., dichloromethane) to convert the terminal amine into a carbamate, where R is alkyl, preferably lower alkyl. Preferably, the inert diluent contains an excess (e.g., about 10 equivalents) of a tertiary amine, such as diisopropylethylamine, to scavenge any acid generated during reaction. Reaction conditions are otherwise conventional as described above. Urea groups are formed at the amino terminus by reaction with an equivalent amount or an excess (e.g., 5 equivalents) of RN=C=O in a suitable inert diluent (e.g., dichloromethane) to convert the terminal amine into a urea (i.e., RNHC(O)NH-) group where R is as defined above. Preferably, the inert diluent contains an excess (e.g., about 10 equivalents) of a tertiary amine, such as diisopropylethylamine. Reaction conditions are otherwise conventional as described above.

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In preparing peptide mimetics wherein the C-terminal carboxyl group is replaced

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by an ester (e.g., -C(O)OR where R is alkyl and preferably lower alkyl), resins used to prepare the peptide acids are employed, and the side chain protected peptide is cleaved with base and the appropriate alcohol, e.g., methanol. Side chain protecting groups are then removed in the usual fashion by treatment with hydrogen fluoride to obtain the desired ester. In preparing peptide mimetics wherein the C-terminal carboxyl group is replaced by the amide -C(O)NR₃R₄, a benzhydrylamine resin is used as the solid support for peptide synthesis, wherein R, and R, are independently alkyl, most preferably lower alkyl. Upon completion of synthesis, hydrogen fluoride treatment to release the peptide from the support results directly in the free peptide amide (i.e., the C-terminus is -C(O)NH₂). Alternatively, use of chloromethylated resin during peptide synthesis coupled with reaction with ammonia to cleave the side chain protected peptide from the support yields the free peptide amide, and reaction with an alkylamine or a dialkylamine yields a side chain protected alkylamide or dialkylamide (i.e., the C-terminus is -C(O)NRR, where R and R₁ are independently alkyl and preferably lower alkyl). Side chain protection is then removed in the usual fashion by treatment with hydrogen fluoride to give the free amides, alkylamides, or dialkylamides.

In another alternative embodiment, the C-terminal carboxyl group or a C-terminal ester can be induced to cyclize by displacement of the -OH or the ester (-OR) of the carboxyl group or ester, respectively, with the N-terminal amino group to form a cyclic peptide. For example, after synthesis and cleavage to give the peptide acid, the free acid is converted in solution to an activated ester by an appropriate carboxyl group activator such as dicyclohexylcarbodiimide (DCC), for example, in methylene chloride (CH₂Cl₂), dimethyl formamide (DMF), or mixtures thereof. The cyclic peptide is then formed by displacement of the activated ester with the N-terminal amine. Cyclization, rather than polymerization, can be enhanced by use of very dilute solutions according to methods well known in the art.

Peptide mimetics as understood in the art and provided by the invention are structurally similar to the paradigm polypeptide comprising the sequence Arg-Xaa-Xaa-Arg (SEQ ID No. 2), but have one or more peptide linkages optionally replaced by a linkage selected from the group consisting of: --CH₂NH--, --CH₂S--, --CH₂CH₂ --, --CH=CH- (in both *cis* and *trans* conformers), --COCH₂--, --CH(OH)CH₂ --, and --CH₂SO--, by methods known in the art and further described in the following references: Spatola,1983, *in* CHEMISTRY AND BIOCHEMISTRY OF AMINO ACIDS, PEPTIDES, AND

PROTEINS, (Weinstein, ed.), Marcel Dekker: New York, p. 267; Spatola, 1983, Peptide Backbone Modifications 1: 3; Morley, 1980, Trends Pharm. Sci. pp. 463-468; Hudson et al., 1979, Int. J. Pept. Prot. Res. 14: 177-185; Spatola et al., 1986, Life Sci. 38: 1243-1249; Hann, 1982, J. Chem. Soc. Perkin Trans. I 307-314; Almquist et al., 1980, J. Med. Chem. 23: 1392-1398; Jennings-White et al., 1982, Tetrahedron Lett. 23: 2533; Szelke et al., 1982, European Patent Application, Publication No. EP045665A; Holladay et al., 1983, Tetrahedron Lett. 24: 4401-4404; and Hruby, 1982, Life Sci. 31: 189-199, each of which is incorporated herein by reference. Such peptide mimetics may have significant advantages over polypeptide embodiments, including, for example: being more economical to produce, having greater chemical stability or enhanced pharmacological properties (such half-life, absorption, potency, efficacy, etc.), reduced antigenicity, and other properties.

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The invention also provides a detailed structural determination of the reactive site loop (RSL) of the PDX protein. The importance of this structure determination, inter alia, relates to the determination that this structure, the RSL, forms a rigid backbone having the positively-charged guanidino residues of each of the Arg residues extending in space away in the same direction from the rest of the PDX protein. This determination results in the capacity to produce peptido-, organo- and chemical mimetics of this portion of the PDX protein structure, said mimetics being capable of inhibiting furin protease activity.

In a second embodiment, the invention provides organic molecules designed to mimic the peptides of the invention by having chemically-similar atoms, moieties or collections thereof in positions analogous to the positions of the atoms, moieties and collections thereof in the Arg-Xaa-Xaa-Arg-comprising peptides of the invention. In a preferred embodiment, these mimetic compounds have the structure:

C(L1-R1)-E-F-G-H-I-J(L2-R2)

wherein "C" is equivalent to the alpha carbon of the first arginine residue, and "J" is equivalent to the alpha carbon of the second arginine residue, in the Arg-Xaa-Xaa-Arg-containing peptides of the invention. Most preferably, "C" and "J" are conformationally hindered as described herein to enable the mimetic to stably adopt the configuration for the (L1-R1) and (L2-R2) substituents present in the Arg-Xaa-Xaa-Arg-containing

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peptides of the invention. "E", "G", and "I" represent planar moieties having dimensions similar to those of a peptide bond (as disclosed in Metzler, 1977, BIOCHEMISTRY: THE CHEMICAL REACTIONS OF LIVING CELLS, Academic Press: New York. p.64). Non-limiting examples of such moieties include vinyl groups and substituted vinyl groups. "F" and "H" are equivalent to the alpha carbons of the two "Xaa" residues in the peptides of the invention, and are preferably conformationally-hindered. for example, by having "H" be restricted in a cyclopentane, cyclopentene, furan. tetrahydrofuran, thiophene, pyrrole, or pyrrolidine ring structure, or by covalent linkage to sterically-hindered groups, such as t-butyl, phenyl, benzyl or substituted phenyl or benzyl groups. The structure represented by E-F-G-H-I is most preferably substantially planar and deviates from this planar structure (for example, by bending, defined herein as flexion above or below the plane defined by E-F-G-H-I) by no more than from about 1 to about 20 degrees, more preferably from about 1 to about 10 degrees, and most preferably by no more than about 5 degrees from said plane. The length of the molecule along the distance between the "C" and "J" components (C-E-F-G-H-I-J) is preferably from about 7.5 to about 11.5 Angstroms, more preferably from about 8.5 to about 10.5 Angstroms, and most preferably about 9.5 Angstroms. R1 and R2 are positivelycharged residues linked to "C" and "J", respectively, by linker groups L1 and L2, respectively, wherein the distance between "C" and L1 and the distance between "J" and L2 is substantially equivalent to the distances between the alpha carbon atoms and the guanidino groups of each of the arginine residues in the Arg-Xaa-Xaa-Arg-containing peptides of the invention. Preferably, R1 and R2 are from about 5 to about 7 Angstroms. more preferably from about 5.6 to about 6.7 Angstroms, and most preferably about 6.2 Angstroms away from their respective alpha carbon equivalents, "C" and "J", and the length of L1 and L2 is chosen to maintain this relative positioning in the mimetic molecule. In addition, R1 and R2 are displaced relative to each other along the longitudinal axis of the molecule to subtend an angle of from about 15 to about 25 degrees, more preferably from about 18 to about 21 degrees, and most preferably about 20 degrees. This arrangement is illustrated in Figure 1B, Left and Right, where the arginine sidechains of both arginine residues in the Arg-Ile-Pro-Arg pharmacophore of the preferred peptide of the invention are shown in orange and red.

The invention provides a pharmacophore for the reactive site loop of the α_1 -antitrypsin variant Portland (SEQ ID No. 1) defined by the sequence Arg₃₅₅-Ile₃₅₆-Pro₃₅₇-

Arg₃₅₈ in this protein. This pharmacophore is represented in Figure 1A, wherein the atom designated "3" is the alpha carbon atom of Arg₃₅₅, and the atom designated "12" is the alpha carbon atom of Arg₃₅₈. These values were derived from the analysis of the crystal structure of the Portland protein as generated by the SYBYL® program (Tripos, Inc., St. Louis, MO); the complete structural information used in these analyses is contained in Appendix A disclosed herewith. The specific portion of these data relating to the sequence Arg₃₅₅-Ile₃₅₆-Pro₃₅₇-Arg₃₅₈ is as follows:

ATOM	2635	N	ARG		-21,632				0 29.44
ATOM	2636	CA	ARG		-23.045				0 28.26
MOTA	. 2637	C	ARG		-23.749				0 30.07
MOTA	2638	0	ARG	355	-24.289				0 31.20
MOTA	2639	CB	ARG	355	-23.609				
MOTA	2640	CG	ARG	355	-22.959				
ATOM	2641	.CD	ARG	355	-23.486		15.145		
ATOM	2642	NE	ARG	355	-22.825	22.976	.15.457		
ATOM	2643	CZ	ARG	355	-23.124	24.073	14.823	1.0	
MOTA	2644	NH	1 ARG	355	-24.026	24.089	13.887	1.0	
ATOM	2645	NH:	2 ARG	355	-22.507	25.176	15.134	1.0	
ATOM	2645	N	ILE	356		15.932	17.265		21.27
MOTA	2647	CA	ILE	356	-24.424	14.667	17.057		19.01
ATOM	2648	C	ILE	356	-25.898	14.821	17.330		26.05
ATOM.	2649	0	ILE	356	-26.268	15.297	18.392		26.38
ATOM	2650	CB	ILE	356	-23.767	13.551	17.907		20.54
ATCM	2651	CG1	ILE	356	-22.307	13.409	17.503		19.5C
MOTA	2652	· cg2	ILE	356	-24.521	12.219	17.649		21.25
ATOM	2653	CD1	ILE	356	-21.595	12.415	18.434		17.49
ATOM	2654	N	PRO	357	-26.758	14.439	16.364		24.53
ATOM	2655	CA	PRO	357	-29.183	14.615	16.534		24.85
ATOM	2656	C	PRO	357	-28.782	13.579	17.447		33.18
MOTA	2657	0	PRO	357	-28.209	12.517	17.631	1.00	
ATOM	2658	CB	PRO	357	-28.715	14.367	15.106		25.84
ATOM	2659	CG	PRO	357	-27.572	13.703	14.306		29.85
ATOM	2660	CD	PRO	357	-26.275	13.877			24.59
MOTA	2661	N	ARG	358	-29.953	13.908	18.026		31.16
ATOM	2662	CA	ARG	358	-30.616	12.936	18.877	_	30.88
MOTA	2663	С	ARG	358	-31.428	12.004	18.024		31.32
ATOM	2664	0	ARG	358	-32.569	12.288	17.695		31.76
MOTA	2665	CB	ARG	358	-31.466	13.637	19.957	1.00	0.00
ATOM	2666	CĢ	ARG	358	-32.415	14.682	19.338	1.00	0.00
ATOM	2667	CD	ARG	358	-33.096	15.460	20.478	1.00	0.00
•				260	-34.540	15.347	20.380	1.00	0.00
ATOM	2668	NE	ARG	358	-35.327	16.130	21.061	1.00	0.00
MOTA	2669	CZ	ARG	358	-34.853	17.034	21.867	1.00	0.00
MOTA	2670		ARG	358	-36.616	16.007	20.933	1.00	0.00
ATOM	2671	NH2	ARG	358	-30.016	20.00.			

wherein "ATOM" indicates the number of the atom in the analyzed sequence, the residue number is shown in column 5, the chemical identity of the residue at each position is shown in column 4, and the analyzed atom is shown in column 3. In column 3, "N" is a peptide nitrogen, "CA" is the alpha carbon, "C" is the peptide carbonyl carbon, "O" is the peptide carbonyl oxygen, "CB", "CG", and "CD" are sidechain methyl or methylene carbon atoms, "NE" is the imino nitrogen of the guanidino group of arginine, "CZ" is the carbon atom of the guanidino group of arginine, and "NH1" and "NH2" are the amino nitrogen atoms of the guanidino group of arginine. Columns 6, 7 and 8 represent the positional information of each atom in the x, y and z axes, respectively. These aspects of the invention are further illustrated in Figure 1B, which shows perspective views of the SYBYL model of the RSL of PDX.

The positional information relating the atoms in this structure is shown in Tables I, II and III:

TABLE I

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Angle (degrees)
31.64
36.93
36.26
35.05
29.95
31.94
35.69
29.22
30.85
30.1
31.75
36.07
35.72
30.48
34.3

TABLE II

Atoms	Distance					
	(Angstroms)					
3-3.5	6.18					
12-12.5	6.1					
3-12	9.48					

TABLE III

Atom	Degrees									
	W .	φ (phi)	ψ (psi)							
3	-178.5	-87	85.29							
6	179.09	82.73	126							
9	-177.2	-76.9	157.9							
12	177.45	-76.9	157.9							

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The complete positional data for the PDX protein is disclosed herein in Appendix A.

Mimetic analogs of the Arg-Xaa-Xaa-Arg-containing peptides of the invention may also be obtained using the principles of conventional or rational drug design (see, Andrews et al., 1990, Proc. Alfred Benzon Symp. 28: 145-165; McPherson, 1990, Eur. J. Biochem. 189:1-24; Hol et al., 1989a, in Molecular Recognition: Chemical and Biochemical Problems, (Roberts, ed.), Royal Society of Chemistry; pp. 84-93; Hol, 1989b, Arzneim-Forsch. 39:1016-1018; Hol, 1986, Agnew Chem. Int. Ed. Engl. 25: 767-778. the disclosures of which are herein incorporated by reference).

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In accordance with the methods of conventional drug design, the desired mimetic molecules are obtained by randomly testing molecules whose structures have an attribute in common with the structure of a "native" Arg-Xaa-Xaa-Arg peptide. The quantitative contribution that results from a change in a particular group of a binding molecule can be determined by measuring the biological activity of the putative mimetic (furin-inhibiting activity) in comparison with the furin-inhibiting activity of the Arg-Xaa-Xaa-Arg-containing peptide. In a preferred embodiment of rational drug design, the mimetic is designed to share an attribute of the most stable three-dimensional conformation of

the Arg-Xaa-Xaa-Arg peptide. Thus, for example, the mimetic may be designed to possess chemical groups that are oriented in a way sufficient to cause ionic, hydrophobic, or van der Waals interactions that are similar to those exhibited by the furin-inhibiting peptides of the invention, as disclosed herein.

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The preferred method for performing rational mimetic design employs a computer system capable of forming a representation of the three-dimensional structure of the Arg-Xaa-Xaa-Arg-containing peptide, such as those exemplified by Hol, 1989a, *ibid.*; Hol, 1989b, *ibid.*; and Hol, 1986, *ibid.* Molecular structures of the peptido-, organo- and chemical mimetics of the peptides of the invention are produced according to those with skill in the art using computer-assisted design programs commercially available in the art. Examples of such programs include SYBYL 6.5®, HQSAR™, and ALCHEMY 2000™ (Tripos); GALAXY™ and AM2000™ (AM Technologies, Inc., San Antonio, TX); CATALYST™ and CERIUS™ (Molecular Simulations, Inc., San Diego, CA); CACHE PRODUCTS™, TSAR™, AMBER™, and CHEM-X™ (Oxford Molecular Products, Oxford, CA) and CHEMBUILDER3D™ (Interactive Simulations, Inc., San Diego, CA).

The peptido-, organo- and chemical mimetics produced using the positional information disclosed herein using, for example, art-recognized molecular modeling programs are produced using conventional chemical synthetic techniques, most preferably designed to accommodate high throughput screening, including combinatorial chemistry methods. Combinatorial methods useful in the production of the peptidoorgano- and chemical mimetics of the invention include phage display arrays, solidphase synthesis and combinatorial chemistry arrays, as provided, for example, by SIDDCO, Tucson, Arizona; Tripos, Inc.; Calbiochem/Novabiochem, San Diego, CA; Symyx Technologies, Inc., Santa Clara, CA; Medichem Research, Inc., Lemont, IL; Pharm-Eco Laboratories, Inc., Bethlehem, PA; or N.V. Organon, Oss, Netherlands. Combinatorial chemistry production of the peptido-, organo- and chemical mimetics of the invention are produced according to methods known in the art, including but not limited to techniques disclosed in Terrett, 1998, COMBINATORIAL CHEMISTRY, Oxford University Press, London; Gallop et al., 1994, "Applications of combinatorial technologies to drug discovery. 1. Background and peptide combinatorial libraries," J. Med. Chem. 37: 1233-51; Gordon et al., 1994, "Applications of combinatorial technologies to drug discovery. 2. Combinatorial organic synthesis, library screening strategies, and future directions," J. Med. Chem. 37: 1385-1401; Look et al., 1996,

Bioorg. Med. Chem. Lett. 6: 707-12; Ruhland et al., 1996, J. Amer. Chem. Soc. 118: 253-4; Gordon et al., 1996, Acc. Chem. Res. 29: 144-54; Thompson & Ellman, 1996, Chem. Rev. 96: 555-600; Fruchtel & Jung, 1996, Angew. Chem. Int. Ed. Engl. 35: 17-42; Pavia, 1995, "The Chemical Generation of Molecular Diversity", Network Science Center, www.netsci.org; Adnan et al., 1995, "Solid Support Combinatorial Chemistry in Lead Discovery and SAR Optimization," Id., Davies and Briant, 1995, "Combinatorial Chemistry Library Design using Pharmacophore Diversity," Id., Pavia, 1996, "Chemically Generated Screening Libraries: Present and Future," Id.; and U.S. Patents, Nos. 5,880,972 to Horlbeck; 5,463,564 to Agrafiotis et al.; 5,331573 to Balaji et al.; and 5,573,905 to Lerner et al.

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The invention also provides antibacterial and antiviral methods. The invention provides methods for blocking endoproteolytic activation of bacterial toxins. Bacterial targets of the antibacterial methods provided by this invention include but are not limited to any bacteria that produces an endoproteolytically-activated toxin, such as diphtheria toxin produced by *Corynebacterium diptheriae*, exotoxin A of *Pseudomonas aurigenosa*, tetanus toxin, the enterotoxins of *Escherichia coli* and *Vibrio cholerae*, protective antigen of *Bacillus anthracis* and the neurotoxin and C2 toxin of *Clostridium botulinum*. Preferred toxins are those that are proteolytically processed at a consensus furin recognition site (-Arg-Xaa-Xaa-Arg-). Preferred embodiments include *Corynebacterium diptheriae*, *Pseudomonas aeruginosa* and *Bacillis anthracis*.

Viral targets of antiviral methods provided include but are not limited to picomaviruses (e.g., poliovirus and rhinovirus); orthomyxovirusus (e.g., influenza virus); paramyxoviruses (e.g., measles virus and mumps virus); coronaviruses; rhabdoviruses (e.g., rabies virus and vesicular stomatitis virus); togaviruses (e.g., Semliki Forest virus and yellow fever virus); bunyaviruses (e.g., California encephalitis virus); arenaviruses (e.g., Lassa fever virus); rubella virus; reoviruses (e.g., Colorado tick fever virus); hepatitis viruses; adenoviruses; herpesviruses (e.g., herpes simplex virus); and oncogenic viruses, including papilloma viruses, RNA tumor viruses, or retroviruses, and lentiviruses (e.g., human immune deficiency virus). The most preferred viruses are the human immunodeficiency viruses (HIV-1 and HIV-2) and human cytomegalovirus (HCMV).

Cells intended to be protected by the methods provided by this invention include but are not limited to human, canine, bovine, murine, leporine, porcine, ovine, simian,

feline, hircine, and equine cells. The preferred cells are human cells. More preferred cells are human T lymphocytes (T cells), and the most preferred human T cells are those human T cells expressing the cell surface antigen CD4.

The methods of the present invention may be used to treat donated human blood or plasma to protect transfusion recipients from viral infection from contaminating virus. The methods of the present invention may be used to treat human semen to protect embryos derived from such semen, and mothers bearing such embryos or impregnated with such semen, from contaminating virus. In a preferred embodiment, the contaminating virus is HIV-1. In another preferred embodiment, the contaminating virus is HCMV.

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The present invention provides methods for inhibiting bacterial infection in a human. The invention also provided for treating a human infected with a bacteria. These methods provided by the invention comprise the step of administering a therapeutically-effective amount of a peptide or peptide mimetic of the invention to the human, most preferably as a pharmaceutical composition comprising a pharmaceutically-acceptable carrier. The invention also provides pharmaceutically acceptable compositions effective for use with the methods provided by the invention comprising the peptides and peptide mimetics of the invention and a pharmaceutically acceptable carrier.

The present invention provides methods for inhibiting viral infection in a human. The invention also provided for treating a human infected with a virus. These methods provided by the invention comprise the step of administering a therapeutically-effective amount of a peptide or peptide mimetic of the invention to the human, most preferably as a pharmaceutical composition comprising a pharmaceutically-acceptable carrier. Preferred viruses of these embodiments of the invention are HIV-1 and HCMV. The invention also provides pharmaceutically acceptable compositions effective for use with the methods provided by the invention comprising the peptides and peptide mimetics of the invention and a pharmaceutically acceptable carrier.

Another embodiment of the present invention includes methods for treating immunosuppression in a human associated with viral infection. Yet another embodiment of the present invention provides a method of prophylaxis for treating a human exposed to infection with a virus, in a particular those directly at risk of infection as a result of intimate contact with humans infected with a virus of tissues or bodily

fluids contaminated by a virus. The preferred virus of these embodiments of the invention is HIV-1. The invention also provides pharmaceutically acceptable compositions effective for use with the methods provided by the invention comprising the peptides and peptide mimetics of the invention and a pharmaceutically acceptable carrier.

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The invention also provides methods for inhibiting proteolytic processing of a biologically active protein or peptide in a cell comprising contacting such cells with the gene therapy delivery system of the invention. The methods of the invention encompass inhibition of proteolytic processing of any biologically active molecule that is proteolytically processed by furin *in vivo* or *in vitro*, including but not limited to peptide hormones, neuropeptides, growth factors, coagulation factors, serum albumin, cell surface receptors, and adhesion molecules. Preferred biologically active proteins are pro-β-nerve growth factor, blood coagulation factor protein Factor IX, pro-von Willibrand factor, complement factor C3 and renin, for alleviation of pathological conditions and disease states in an animal, preferably a human, associated with over-expression, over-production or otherwise inappropriate synthesis of such biologically-active proteins.

Preparation of pharmaceutically acceptable compositions provided by the present invention can be prepared using methods well know to those with skill in the art. Any of the common pharmaceutical-acceptable carriers such as sterile saline solution, plasma, etc., can be utilized for preparing the pharmaceutical compositions provided by the invention. Routes of administration include but are not limited to oral, nasal (including inhalation into the lungs), intravenous, parenteral, rectal, optical, aural and transdermal. The pharmaceutical compositions of the invention may be administered intravenously in any conventional medium for intravenous injection such as a aqueous saline medium, or in blood plasma medium. Such medium may also contain conventional pharmaceutical adjunct materials such as, for example, pharmaceutically acceptable salts to adjust the osmotic pressure, buffers, preservatives and the like. Among the preferred media are normal saline and plasma.

Formulations may further include one or more diluents, fillers, emulsifiers, preservatives, buffers, excipients, and the like, and may be provided in such forms as liquids, powders, emulsions, suppositories, liposomes, transdermal patches and tablets, for example. The agents of the present invention can be formulated according to known

methods to prepare pharmaceutically useful compositions, whereby these materials, or their functional derivatives, are combined in admixture with a pharmaceutically acceptable carrier vehicle. Suitable vehicles and their formulation, inclusive of other human proteins, e.g., human serum albumin, are described, for example, in REMINGTON'S PHARMACEUTICAL SCIENCES (1980, 16th ed., Osol, ed., Mack Press:Easton PA).

Additional pharmaceutical methods may be employed to control the duration of action. Control release preparations may be achieved through the use of polymers to complex or absorb a mimetic of the invention. Such controlled delivery may be exercised by selecting appropriate macromolecules (for example polyesters, polyamino acids, polyvinyl pyrrolidone, ethylenevinylacetate, methylcellulose, carboxymethylcellulose, or protamine, sulfate) and the concentration of macromolecules as well as the methods of incorporation in order to control release. Another possible method to control the duration of action by controlled release preparations is to incorporate a peptide or peptide mimetic of the invention into particles of a polymeric material such as polyesters, polyamino acids, hydrogels, poly(lactic acid) or ethylene vinylacetate copolymers. Alternatively, instead of incorporating these agents into polymeric particles, it is possible to entrap these materials in microcapsules prepared, for example, by coacervation techniques or by interfacial polymerization, for example, hydroxymethylcellulose or gelatine-microcapsules and poly(methylmethacylate) microcapsules, respectively, or in colloidal drug delivery systems, for example, liposomes, albumin microspheres, microemulsions, nanoparticles, and nanocapsules or in macroemulsions. Such techniques are disclosed in REMINGTON'S PHARMACEUTICAL Sciences (1980).

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Compositions for parenteral administration include sterile aqueous or non-aqueous solutions, suspensions, and emulsions. Examples of non-aqueous solvents are propylene glycol, polyethylene glycol, vegetable oils such as olive oil, and injectable organic esters such as ethyl oleate. Carriers, adjuncts or occlusive dressings can be used to increase tissue permeability and enhance antigen absorption. Liquid dosage forms for oral administration may generally comprise a liposome solution containing the liquid dosage form. Suitable forms for suspending liposomes include emulsions, suspensions, solutions, syrups, and elixirs containing inert diluents commonly used in the art, such as purified water. Besides the inert diluents, such compositions can also include wetting

agents, emulsifying and suspending agents, or sweetening, flavoring, coloring or perfuming agents.

A composition is said to be "pharmacologically acceptable" if its administration can be tolerated by a recipient patient. Such an agent is said to be administered in a "therapeutically effective amount" if the amount administered is physiologically significant. An agent is physiologically significant if its presence results in a detectable change in the physiology of a recipient patient.

Generally, the dosage needed to provide an effective amount of the composition will vary depending upon such factors as the recipient's age, condition, gender, and extent of disease, if any, and other variables which can be adjusted by one of ordinary skill in the art. The pharmaceutical compositions and medicaments of the invention are preferably in the form of a unit dose in solid, semi-solid and liquid dosage forms such as tablets, pills, powders, liquid solutions or suspensions, and injectable and infusible solutions. Effective dosage ranges from about 100 µg/kg to about 10 mg/kg of body weight are contemplated.

The Examples which follow are illustrative of specific embodiments of the invention, and various uses thereof. They set forth for explanatory purposes only, and are not to be taken as limiting the invention.

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EXAMPLE 1

Furin Inhibition Assay

In order to assess the biological activity of the mimetics of the invention, a furin inhibition assay was developed as follows.

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A model system was prepared for assaying furin-catalyzed maturation of *Pseudomonas aeruginosa* exotoxion A (PEA), as shown in Figures 2, 3 and 4. *P. aeruginosa* pro-exotoxion A is cleaved at the sequence Arg-Gln-Pro-Arg₂₇₉ in the II + Ib subunit of the toxin in endosomes of infected cells (Figure 2). In the assay, illustrated in Figure 3, A7 cells were incubated in the presence and absence of test PDX mimetics of the invention for 1h under cell culture conditions. The media containing the mimetic was then exchanged for fresh media containing a growth-inhibitory amount of pro-PEA, and incubated under cell culture conditions for 6h. Thereafter, the cells were metabolically labeled with ³⁵S-labeled methionine and/or cysteine for 30min, and

cellular proteins precipitated with trichloroacetic acid. PDX itself or an Arg-Xaa-Xaa-Arg-containing peptide was used as a positive control, and α_1 -antitrypsin Pittsburgh (PIT; SEQ ID No. 6) was used as a negative control in these assays.

The results of a standardized test assay showing the difference in protein synthesis in the presence of PDX or PIT is shown in Figure 4. Preincubation of A7 cells in the presence of increasing concentrations of PDX resulted in increasing levels of protein synthesis in the presence of PEA compared with cells incubated without PDX. In contrast, little or no protective effect was observed by incubating A7 cells with PIT at any concentration tested. These results indicated that the assay was effective in detecting PDX-mediated inhibition of furin-catalyzed maturation of pro-PEA.

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An alternative assay was used to demonstrate inhibition of furin-mediated maturation of human cytomegalovirus (HCMV) glycoprotein gB. This assay is illustrated in Figures 2, and 5 through 8. HCMV glycoprotein pro-gB is cleaved at the sequence Arg-Thr-Lys-Arg₄₆₅ in the pro-protein, yielding gp110 and gp55, linked by a disulfide bond, as shown in Figure 5. In the assay, illustrated in Figure 6, U373 cells were infected with HCMV Towne at a multiplicity of infection of about 0.1. After infection, the putative inhibitor was added and incubated with the infected cells for 5 days under cell culture conditions. Cell extracts were then prepared and used to infect a naive culture of human foreskin fibroblasts (HFF), which were incubated in the absence of inhibitor for 7 days. These cells were immobilized under agar using conventional techniques, and the number of infected viral plaques determined by counting. Foscarnet, a phosphate group analog and known HCMV inhibitor, was assayed in parallel as a positive control.

The results of a standardized test assay using PDX and foscarnet are shown in Figure 7. PDX has an ED₅₀ that is about ten-fold lower than foscarnet for plaque formation, illustrating its enhanced inhibitory capacity. These results indicated that the assay was effective in detecting PDX-mediated inhibition of furin-catalyzed maturation of HCMV glycoprotein gB.

In an additional or alternative embodiment of this assay, parallel cultures of infected U373 cells were grown in the presence of inhibitor and cellular proteins isolated after 5 days of infected cell growth. Sodium dodecyl sulfate/polyacrylamide gel electrophoresis (SDS/PAGE) was performed on the cellular protein extract, followed by Western blotting and hybridization with an immunological reagent specific for

glycoprotein gB, all techniques performed as described in Sambrook *et al.* (1989, MOLECULAR CLONING: A LABORATORY MANUAL, CSPLP: New York). In these assays PDX itself or an Arg-Xaa-Xaa-Arg-containing peptide was used as a positive control, and α_1 -antitrypsin Pittsburgh (PIT; SEQ ID No. 6) was used as a negative control.

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The results of a standardized test assay showing the difference in HCMV glycoprotein gB maturation in the presence of PDX or PIT is shown in Figure 8. In the absence of either PDX or PIT, infected cells were observed to contain predominantly gp110 and gp55, separated by treatment with β -mercaptoethanol prior to SDS-PAGE analysis (lane labeled gB/Wt in the Figure). A similar level of cleavage was seen for cells incubated with the PIT variant of α_1 -antitrypsin (lane labeled gB/PIT) In contrast, the predominant band observed in the cell extract from infected cells incubated in the presence of PDX was pro-gB (lane labeled gB/PDX). These results indicated that the assay was effective in detecting PDX-mediated inhibition of furin-catalyzed maturation of HCMV glycoprotein gB.

These assays are used to characterize the furin inhibitory capacity of the mimetic compounds of the invention. Preferably, cells are incubated in varying concentrations of the mimetic, in parallel with a standardized concentration of PDX or an Arg-Xaa-Xaa-Arg-containing peptide. Furin inhibitory capacity of putatuve mimetics of the invention are characterized by quantitative comparisons of the extent of furin inhibition, measured as described herein by ED₅₀ of plaque formation, percent protein synthesis, or K_1 of furin activity, to PDX or Arg-Xaa-Xaa-Arg-containing peptides of the invention.

It should be understood that the foregoing disclosure emphasizes certain specific embodiments of the invention and that all modifications or alternatives equivalent thereto are within the spirit and scope of the invention as set forth in the appended claims. The disclosure of all patents, patent applications and publications are hereby incorporated by reference herein in their entirety.

APPENDIX A

HEADER:		PRO	YTEIN									18-1	LAR-9	Ω	
COMPND:			hal-PI	x:	_							•••	5 -2\-,		
AUTHOR:		•	ncois		GEN	ERAT	ED E	Y SY	BYL.	A P	RODU	וכד כ	F TR	TPOS	:
ASSOCIA	TES,				•				•						
SEQRES	1	3	72 PH	E ASN	LYS	İLE	THR	PRO	ASN	LEU	ALA	GLU	PHE	ALA	PHE
SEQRES	2	. 3		R LEU											
SEQRES	. 3	3		N ÌLE											
SEQRES	4	3	72 AL	A MET	LEU	SER	LEU	GLY	THR	LYS	ALA	ASP	THR	HIS	ASP
SEQRES	5	3		U ILE											
SEQRES	6	3		o GLU											
SEQRES	7			G THR											
SEQRES	8			R GLY											
SEQRES	9			L ASP											
SEQRES	10			R GLU											
SEQRES	11	-		A LYS											
SEQRES	12	_		N GLY											
SEQRES	13			P THR											
SEQRES	14			LYS											
SEQRES				J GLU											
SEQRES	16 17			PRO											
SEQRES	18			CYS											
SEQRES	19			J GLY											
SEQRES	20			ILE											
SEORES	21			ALA											
SEORES	22	37		TYR											
SEQRES	23	3 7		LYS											
SEQRES	24	37		GLU											
SEQRES	25	37		ALA											
SEQRES	26	37		GLY											
SEQRES	27	37		PRO											
SEQRES	28	37		ILE							PRO	LEU	PHE :	MET (GLY -
	29	37	2 LYS	VAĹ	VAL	ASN	PRO			LYS		•	•		•
MOTA	1	N	PHE	23			833		226	-6.0	590		0 23		
MOTA	2	CA	PHE	23		-5.		16.		-8.:			0 22		
MOTA	3	С	PHE	23		-4.			813	-8.5			0 27		
ATOM	4	0	PHE	23			316	14.		-7.6			0 30		
ATOM ATOM	5. 6		PHE	23		-4.		17.2		-B.6			0 23		
ATOM	7	CG	PHE	23		-3.		17.3		-7.9			24.		
ATOM	. 8	CD2	PHE PHE	23 23		-3.3 -2.3		18.0		-6.7			27.		
ATOM	9		PHE	23		-1.9		16.6		-8.5) 28.) 31.	-	•
ATOM	10		PHE	23			991			-6.1 -7.8) 31.) 29.		
ATOM	11	cz	PHE	23		-0.8		17.4		-5.6			29.		
ATOM	12	N	ASN	24		-5.0		14.4		-9.7			21.		
ATOM	13	CA	ASN -	24		-4.5				10.2			22.		
ATOM	14	C	ASN	24		-3.7				11.6			27.		
MOTA	15	0	AŚN	24	•	-4.0				12.2			27.		
MOTA	16	СВ	ASN	24		-5.7				10.5			22.		
ATOM '	17	CG	ASN	24		-6.8		12.4		-9.5			53.		
MOTA	18	OD1	ASN	24		-6.6	82	12.1		-8.3			51.		
MOTA	19	ND2		24		-7.9		13.0	68	-9.9			50.		
MOTA	20	N	LYS	25		-2.8	88	12.5	09 -	11.9	81	1.00	23.	21	
MOTA	21	CA	LYS	25	`	-2.1	12	12.6	64 -	13.2	16	1.00	23.	36	

ATOM	22	С	LYS	25	-1.863	11.278 -13.83	
MOTA	23	0	LYS	25	-2.147		
MOTA	24	. CB	LYS	25	-0.780	13.350 -12.86	
MOTA	25	CG	LYS	25	-0.046	•	
MOTA	26	CD	LYS	25	1.032	14.953 -13.43	
MOTA	27	CE	LYS	25	2.011	15.488 -14.46	
MOTA	28	NZ	LYS	25	3.041	16.381 -13.84	
MOTA	29	N	ILE	26	-1.364	10.360 -13.00	
ATOM	30	CX	ILE	26	-1.080	8.977 -13.39	
ATOM	31	C	ILE	26	-1.881	8.077 -12.45	
ATOM	32	0	ILE	26	-1.928	6.862 -12.636	
ATOM	33	CB	ILE	26	0.431	8.626 -13.231	
MOTA	34	CG1		26	0.800	8.525 -11.759	•
ATOM	35	ÇGZ		26	1.292	9.669 -13.907	
ATOM	36	CD1		26	2.225	8.160 -11.516	
ATOM	37	N	THR	27	-2.535	8.697 -11:.470	
MOTA	38	CA	THR	27	-3.312	7.978 -10.472	
ATOM	39	С	THR	27	-4.328	7.029 -11.048	
MOTA	40	0	THR		-4.554	5.960 -10.482	
ATOM	41	CB	THR	27	-4.013 -5.184	8.924 -9.507 9.463 -10.134	•
ATOM ATOM	42 43	OG1 CG2		27 27	-3.184	10.036 -9.116	
ATOM	44	N	THR PRO	28	-4.995	7.414 -12.147	
ATOM	45	CA	PRO	28	-5.951	6.424 -12.631	
ATOM	46	C	PRO	28	-5.213	5.159 -13.050	
ATOM	47	0	PRO	28	-5.700	4.044 -12.837	•
ATOM	48	СВ	PRO	28	-6.672	7.139 -13.782	
ATOM	49	CG	PRO	28	-5.805	8.336 -14.120	
ATOM	50	CD	PRO	28	-5.158	8.716 -12.819	
ATOM	51	N	ASN	29	-3.978	5.350 -13.515	
ATOM	52	CA	ASN	29	-3.127	4:249 -13.957	
ATOM	53	С	ASN	2.9	-2.694	3.448 -12.733	1.00 21.94
ATOM	54	0	ASN	29	-2.648	2.223 -12.768	1.00 23.88
ATOM	55	СВ	ASN	29	-1.899	4.770 -14.725	1.00 19.53
MOTA	56	CG	ASN	29	-2.261	5.486 -16.027	1.00 48.12
ATOM	57	OD1	ASN	29	-2.498	4.852 -17.054	1.00 41.88
ATOM	58	ND2	ASN	29	-2.252	6.819 -15.999	1.00 43.32
ATOM	59	N	LEU	30	-2.441	4.136 -11.627	1.00 16.53
ATOM	60	CA	LEU	30	-2.033	3.466 -10.392	1.00 16.25
MOTA	61	C	LEU	30 -	-3.195	2.744 -9.735	1.00 22.44
MOTA	62	0	LEU	30	-3.013	1.679 -9.147	1.00 23.44
ATOM	63	CB	LEU	30	-1.405	4.464 -9.402	1.00 16.22
ATOM	64	CG	LEU	30	-0.017	5.008 -9.772	1.00 20.92
ATOM	65		LEU	30	0.443	6.007 -8.746	1.00 20.56
ATOM	66		LEU	30	0.989	3.875 -9.911	1.00 22.13
ATOM	67	N	ALA	31	-4.390	3.311 -9.873	1.00 19.06
ATOM	68	CA	ALA	31	-5.618	2.752 -9.301	1.00 17.69
ATOM	69	C	ALA	31	-5.914	1.378 -9.880 0.387 -9.152	1.00 20.90
ATOM	70	0	ALA	31	-6.024 -6.790		1.00 18.85 1.00 18.32
ATOM	71	CB	ALA	31	-6.790 -6.041	3.681 -9.570 1.328 -11.198	1.00 18.32
ATOM ATOM	72 73	N CA	GLU	32 32	-6.316	0.081 -11.879	1.00 20.59
ATOM	74	C	GLU	32	-5.190	-0.921 -11.671	1.00 24.22
ATOM	75	0	GLU	. 32	-5.393	-2.135 -11.765	1.00 25.34
~100	, ,	•	320		- 3.333		

		-											
ATOM	. 7	6 C	B GLU	32		-6.6	510	. 0.34	11 -13.3	58 1	.00	22.3	
ATOM	7	7 C	G - GLU	32		-7.7			1 -13.5			31.7	
ATOH	71	8 C	D GLU	32		-9.0	57		9 -12.8			49.10	
ATOM	.79	0	E1 GLU	32		-9.1	77		6 -12.3			28.8	
HOTA	80	0	E2 GLU	32		-9.9	77		0 -12.8			50.57	
MOTA	. 81	N	PHE	33		-4.0	11		4 -11.3			18.26	
ATOM	82	. C	A PHE	- 33		-2.8	56		4 -11.0			16.77	
ATOM	-83	C	PHE	33		-3.0	87	-1.88				18.47	
ATOM	84	0	PHE	33	•	2.9	28	-3.09				14.96	
ATOM	. 85		PHE	33		-1.5	73	-0.40	6 -11.00			17.37	
ATOM	86			33		-0.3	97	-1.09	0 -10.43	9 1.	00	17.39	
ATOM	-87		I PHE	33		0.1	30	-2.24	5 -10.99	8 1.	00	21.30	
MOTA	88		2 PHE	33		0.1	61	-0.59			00	19.00	
ATOM			1 PHE			1.20	03	-2.90	3 -10.39		00	22.86	
ATOH	90		2 PHE	33	-	1.2		-1.242				21.58	
ATOH	91			33		1,75		-2.399	-9.22			20.43	
ATOH	92	N	λLA	34		-3.56		-1.066				16.77	
ATOM	93	CA		34		-3.86			-7.37			15.77	
MOTA	94	C	ALA	34	•	-4.93		-2.562				21.38	
ATOM	95	0	ALA	34		-4.84		-3.548				23.09	
ATOM ATOM	96	CB		34		-4.31	-	-0.311				15.65	
ATOM	97	N CA		35		-5.94		-2.381				7.88	
ATOM	99	C	PHE Phe	35 [.]		-7.03		-3.350				6.95	
ATOM	100	0	PHE	35 35		-6.57		-4.663				0.50	
ATOM	101	CB	PHE	35		-6.89 -8.19		-5.740				0.02	
ATOM	102	CG	PHE	35		-8.82		-2.771 -1.551				8.28	
ATOM	103		PHE	35	-	-8.85		-1.392	-8.479 -7.097			9.41	
ATOM	104		PHE	35		-9.38		-0.561	-9.269			2.41 1.84	
ATOM	105		PHE	35		-9.43		-0.264	-6.522			5.39	
ATOM	106		PHE	35		-9.97		0.568				2.84	
ATOM	107	CZ	PHE	35		-9.99		0.715	-7.329			2.70	
MOTA	108	N	SER	36	-	-5.76		4.558	-9.959			B.62	
ATOM	109	CA.	SER	36		-5.232			-10.659			B.48	
ATOM	. 110	C	SER	36		-4.386		6.509	-9.685			9.66	
ATOM	111	0	SER	36		-4.593		7.701	-9.491			7.74	
ATOM	112	CB	SER	36		-4.387	' -	5.258	-11.853			0.88	
ATOM	113	OG:	SER	36		-4.134	-	6.331	-12.735			.56	
MOTA	114	N	LEU	37		-3.499		5.811	-8.998	1.00	14	. 84	
MOTA	115	CA	LEU	37		-2.626		6.459	-8.037	1.00	15	.38	
ATOH	116	C	LEU	37		-3.417		7.009	-6.844	1.00	20	.75	
ATOM	117	0	LEU	37		-3.014		7.991	-6.228	1.00			
HOTA	118	CB	LEU	37		-1.536		5.483	-7.586	1.00			
ATOH	119	CG	LEU	37		-0.433		5.990	-6.656	1.00			
ntom Nota	120		LEU	37		0.354		7.114	-7.294	1.00			
ATOH	121	CD2		37		0.480		4.836	-6.336	1.00			
ATOM ATOM	122 123	N CA	TYR	38		-4.602			-6.613	1.00			
NTOH		CA	TYR TYR	38 38		-5.457		6.869	-5.510	1.00			
NTOH		0		38		-6.204		8.160	-5.836	1.00			
NOT		CB	TYR TYR	38 38		-6.054		9.175	-5.148	1.00			
MOTA		CG	TYR	38		-6.443 -7.398		5.751	-5.153	1.00			
TOM			TYR	38		-7.398 -7.050		5.137	-4.054	1.00			
TOH		CD2		38		-8.616		.991	-2.715	1.00			
			•••	70		-0.010	- 0	.744	-4.352	1.00	19.	. 23	

ATOM	13	0 C	E1 TYR	38	-7.890 -6.453 -1.704 1.00 17.18
ATOM	13	1 C	E2 TYR	38	-9.453 -7.203 -3.350 1.00 20.11
ATOM	13	2 C	Z TYR	38	-9.085 -7.058 -2.037 1.00 24.15
MOTA	13	3 01	H TYR	- 38	-9.915 -7.535 -1.063 1.00 26.53
MOTA	13	N	ARG	39	-6.979 -8.121 -6.913 1.00 17.51
ATOM	139	S C	A ARG	39	-7.761 -9.263 -7.383 1.00 17.33
ATOM	136	s c	ARG	. 39	-6.856 -10.469 -7.617 1.00 19.96
ATOM	137	7 0	ARG	39	-7.296 -11.617 -7.538 1.00 19.48
ATOM	138	CE	ARG	39	-8.526 -8.870 -8.656 1.00 17.17
ATOM	139) cc	ARG	39	-9.451 -7.667 -8.419 1.00 24.64
ATOM	140	CI	ARG	39	-10.224 -7.239 -9.648 1.00 34.67
ATOM	141	. NE	ARG	39	-11.117 -6.112 -9.378 1.00 44.73
MOTA	142	CZ	ARG	39	-10.927 -4.885 -9.849 1.00 66.12
ATOM	143		1 ARG	39	-9.879 -4.622 -10.616 1.00 58.52
ATOM	144	NH	2 ARG	39	-11.786 -3.920 -9.552 1.00 57.78
ATOM	145		GLN	40	-5.581 -10.172 -7.8331.00 16.43
ATOM	146		GLN	40	-4.525 -11.145 -8.058 1.00 17.01
ATOM	147		GLN	40	-4.296 -11.946 -6.780 1.00 24.64
ATOM	148		GLN	40	-4.377 -13.176 -6.757 1.00 26.03
ATOM	149			40	-3.242 -10.379 -8.399 1.00 18.53
ATOM	150			40	-2.028 -11.234 -8.646 1.00 32.27
ATOM	151	CD		40	-2.196 -12.108 -9.853 1.00 65.70
ATOM	152		1 GLN	40	-2.982 -11.814 -10.756 1.00 68.30
MOTA	153		2 GLN	40	-1.469 -13.194 -9.876 1.00 60.66
MOTA	154	N	LEU		-4.036 -11.213 -5.711 1.00 20.30
MOTA	155	CA	LEU	41	-3.769 -11.784 -4.412 1.00 19.36
ATOM	156	С	LEU	41	-5.054 -12.277 -3.745 1.00 25.88
MOTA	157	0	LEU	41	-5.039 -13.298 -3.048 1.00 26.53
ATOM	158	CB		41	-3.067 -10.724 -3.540 1.00 18.71
ATOM ATOM	159 160	CG	LEU LEU	41	-1.757 -10.130 -4.089 1.00 23.44
ATOM	161		LEU .	41 41	-1.403 -8.856 -3.362 1.00 22.82
ATOM	162	N N	ALA	42	-0.631 -11.142 -3.986 1.00 26.55 -6.175 -11.626 -4.053 1.00 21.70
ATOM	163	CA	ALA	42	· · · · · · · · · · · · · · · · · · ·
ATOM	164	c	ALA	42	
ATOM	165	0	ALA	42	-8.019 -13.311 -3.903 1.00 25.53 -8.771 -13.955 -3.176 1.00 26.06
ATOM	166	СВ	ALA	42	-8.502 -10.856 -3.683 1.00 20.25
MOTA	167	N	HIS	43	-7.688 -13.705 -5.130 1.00 20.57
ATOM	168	CA	HIS	43	-8.147 -14.981 -5.657 1.00 20.04
ATOM	169	C	HIS	43	-7.203 -16.082 -5.208 1.00 24.10
ATOM	170	0	HIS	43	-7.619 -17.009 -4.509 1.00 23.78
ATOM	171	CB	HIS	43	-8.237 -14.960 -7.186 1.00 21.07
ATOM	172	CG	HIS	43	-9.585 -14.594 -7.708 1.00 25.01
ATOM	173	ND1	HIS	43	-9.950 -13.285 -7.989 1.00 27.71
MOTA	174	CD2	HIS	43	-10.683 -15.348 -7.974 1.00 26.92
MOTA	175	CE1	HIS	43	-11.203 -13.258 -8.411 1.00 27.25
	176	NE2	HIS	43	-11.661 -14.495 -8.418 1.00 27.36
MOTA	177	N	GLN	44	-5.941 -15.956 -5.615 1.00 20.85
ATOM	178	CA	GLN	44	-4.877 -16.901 -5.276 1.00 21.40
ATOM	179	C	GLN	44	-4.960 -17.227 -3.797 1.00 27.47
ATOM	180	0	GLN	44	-5.420 -18.303 -3.403 1.00 27.52
MOTA	181	CB	GLN	44 .	-3.518 -16.267 -5.590 1.00 23.51
MOTA	182	CG	GLN	44	-2.319 -16.941 -4.935 1.00 67.92
MOTA	183	CD	GLN	44	-1.087 -16.070 -4.998 1.00109.81

ATOM	18	4 0	E1 GL	1 44	-1.07	8 -15.03	1 -5.66	2 1.00109.42
ATOM	18	5 N	E2 GLA	44		516.46		
ATOH	18	6 N	SEF	45		5 -16.25		
ATOM	18	7 C	A SER	45		0 -16.36		
MOTA	18	8 C	SER	45	-6.05	7 -16.28		
ATOM	18	9 0	SER	45	-6.78	7 -15.49	1 -1.800	
MOTA	190	0 C	B SER	45	-3.78	9 -15.194	-0.941	
ATOM	19:		G SER		-3.34	3 -15.506	0.359	
MOTA	192					7 -17.107		1.00 27.55
MOTA	19					7 -17.130		
ATOM	198					-16.608		1.00 28.59
ATOM	195					2 -15.772		
ATOM	196					-18.519		
MOTA	197			46		-19.636		
MOTA	198		1 ASN	46		-19.544	=	
ATOM	199		2 ASN	46		-20.708		
ATOM	200		SER	47		-17.107		
ATOM	201			47		-16.653		
ATOM	202		SER	47		-15.935		1.00 27.21
ATOM ATOM	203 204		SER	47		-16.296		1.00 27.54
ATOM	205			47 47		-17.824	4.742	1.00 24.67
ATOM	205			4.8		-17.784 -14.889	4.892	1.00 37.40
ATOM	207	CA		48		-14.126		1.00 24.53
ATOM	208	c	THR	. 48		-12.602	4.410	1.00 24.98
ATOM	209	ō	THR	48		-12.102	3.477	1.00 29.19 1.00 32.29
ATOM	210	СВ		48		-14.482	3.564	1.00 32.29
MOTA	211	OG		48		-15.851	3.794	1.00 37.64
ATOM	212	CG		48		-13.591	3.903	1.00 37.25
ATOM	213	N	ASN	49		-11.874		1.00 21.11
ATOH	214	CA	ASN	49		-10.413	5.342	1.00 20.02
MOTA	215	C	ASN	49		-10.014	4.197	1.00 25.63
MOTA	216	0	ASN	49	-3.078	-10.599	4.023	1.00 28.91
ATOM,	217	CB	ASN	49	-4.565	-9.775	6.645	1.00 23.44
ATOM	218	CG	ASN	49	-5.366	-10.201	7.851	1.00 57.54
MOTA	219		LASN	49	-6.599	-10.128	7.845	1.00 44.13
MOTA	220		2 ASN	49		-10.672	8.888	1.00 54.42
ATOM	221	N	ILE	50	-4.588	-9.067	3.383	1.00 19.43
MOTA	222	CA	ILE	50-	-3.795	-8.628	2.247	1.00 17.16
ATOM	223	C	ILE	50	-3.352	-7.206	2.512	1.00 19.43
atom Atom	224	0	ILE	50	-4.120	-6.415	3.043	1.00 20.29
ATOM	225 226		ILE	50		-8.692	0.942	1.00 19.64
ATOM	227		ILE	50 50	-5.152 -3.804		0.736	1.00 19.17
ATOM	228		ILE	50	-5.804 -6.136	-8.221	-0.252	1.00 19.61
ATOM	229	N	LEU	51	-2.105	-10.249 -6.885	-0.374	1.00 11.82
MOTA	230	CA	LEU	51	-1.631	-5.526	2.194	1.00 15.23
ATOM	231	C	LEU	51	-0.471	-5.202	2.401 1.491	1.00 15.02 1.00 20.37
MOTA	232	ō	LEU	51	0.576	-5.833		1.00 20.37
MOTA	233	CB	LEU	51	-1.217	-5.303		1.00 21.91
MOTA	234	CG	LEU	51	-0.860	-3.870		1.00 19.98
MOTA	235		LEU	51	-2.033	-2.949		1.00 19.60
MOTA	236		LEU	51	-0.437	-3.830		1.00 20.30
MOTA	237	N	PHE	52	-0.666	-4.247		1.00 15.96
								

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HOTA	238	CA	PHE	52	0.399	-3.844	-0.303	1.00 15.36
MOTA	239	C	PHE	52	0.270	-2.368	-0.613	
MOTA	240	0	PHE	52	-0.803	-1.788	-0.458	1.00 19.47
MOTA	241	СВ	PHE	52	0.390	-4.676	-1.595	1.00 16.46
MOTA	242	CG	PHE	52	-0.890	-4.566	-2.394	1.00 16.66
ATOM	243		1 PHE		-1.957	-5.430	-2.155	1.00 17.01
ATOM	244		2 PHE	52	-1.014	-3.624	-3.410	1.00 16.97
ATOM	245		1 PHE	52	-3.108	-5.356		1.00 18.40
MOTA	246		2 PHE	52	-2.165	-3.552	-4.166	1.00 16.88
MOTA	247	CZ	PHE	52	-3.210	-4.419	-3.917	1.00 15.91
MOTA	248	N	SER	53	1.383	-1.761	-1.006	1.00 14.83
ATOM	249	CA	SER	53	1.391	-0.357	-1.365	1.00 14.34
ATOM	250	C	SER	53	1.428	-0.278	-2.891	1.00 21.85
MOTA	251	0	SER		2.412	-0.704	-3.515	1.00 22.16
ATOH	252	CB	SER	53	2.609	0.332	-0.738	1.00 13.76
ATOM	253	⊙G	SER	53	2.700	1.710	-1.076 -3.519	1.00 14.03
MOTA	254	N	PRO	54	0.317	0.151		1.00 19.23
ATOM	255	CA	PRO	54	0.307	1.295	-4.977 -5.408	
ATOM	256	C.	PRO	54	1.308		-6.443	1.00 21.51
ATOM	257	0	PRO	54	1.957	0.670	-5.276	1.00 22.63
MOTA	258	CB	PRO	54	-1.129 -1.889	-0.022	-4.200	1.00 18.91 1.00 23.13
ATOM	259	CD	PRO	54 54		0.321	-2.994	1.00 23.13
ATOM ATOM	260 261	N ·	PRO VAL	54 55	-1.052 1.539	2.243	-4.501	1.00 19.89
ATOM	262	CA:	VAL	55 55	2.461	3.348	-4.717	1.00 17.72
ATOM	263		VAL	55 55	3.930	2.952	-4.583	1.00 17.43
ATOM	264	0	VAL	55 55	4.702	3.173	-5.511	1.00 23.27
MOTA	265	CB	VAL	.55	2.158	4.528	-3.769	1.00 20.90
ATOM	266		VAL	55	3.299	5.529	-3.784	1.00 20.30
MOTA	267		VAL	55	0.882	5.220	-4.199	1.00 20.16
ATOM	268	N N	SER	56	4.316	2.362	-3.450	1.00 21.09
ATOM	269	CA	SER	56	5.708	1.971	-3.240	1.00 21:01
ATOM	270	C	SER	. 56	6.175	1.003	-4.318	1.00 22.56
ATOM	271	ō	SER	56	7.256	1.172	-4.891	1.00 22.87
ATOM	272	СВ	SER	56	5.942	1.402	-1.822	1.00 23.93
ATOM	273	OG	SER	56	5.405	0.099	-1.647	1.00 27.29
MOTA	274	N	ILE	57	5.312	0.054	-4.664	1.00 15.52
ATOM	275	CA	ILE	57	5.654	-0.936	-5.680	1.00 14.94
MOTA	276	С	ILE	57	5.902	-0.298	-7.053	1.00 17.44
ATOM	277	0	ILE	57	6.986	-0.425	-7.615	1.00 16.01
MOTA	278	CB	ILE	57	4.562	-2.017	-5.806	1.00 18.06
MOTA	279	CG1	ILE	·57	4.511	-2.876	-4.556	1.00 17.03
MOTA	280		ILE	57	4.844	-2.929	-6.974	1.00 20.11
MOTA	281	CD1	ILE	57	3.466	-3.942	-4.631	1.00 17.46
ATOM	282	N	ALA	58	4.894	0.394	-7.570	1.00 14.71
MOTA	283	CA	ALA	58	4.959	1.047	-8.866	1.00 15.31
ATOM	284	С	ALA	58	6.147	1.987	-8.999	1.00 22.96
ATOM	285	0 .	ALA	58	6.987	1.814	-9.885	1.00 25.76
MOTA	286	CB	ALA	58	3.670	1.790	-9.134	1.00 15.90
MOTA	287	N	THR	59	6.203	2.990	-8.131	1.00 16.71
ATOM	288	CA	THR	5 ġ	7.287	3.959	-8.132	1.00 14.91
MOTA	289	C	THR	59	8.656	3.253	-8.180	1.00 19.37
ATOM	290	0	THR	59	9.585	3.726	-8.835	1.00 17.29
ATOM	291	CB	THR	59	7.168	4.863	-6.887	1.00 11.03
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ATOM			THR	59	5.843	5.400	-6.830	
MOTA	293	CG2	THR	59	8.155	6.014	-6.946	1.00 4.73
MOTA	294	N	ALA	60 .	8.711	2.055	-7.604	1.00 17.18
ATOM-	295	CA.		60	9.922	1.246	-7.555	1.00 16.17
MOTA	. 296	C	YLA	60	10.247	0.670	-8.924	1.00 17.24
ATOM	297	0	ALA	60	11.380	0.744	-9.384	1.00 17.12
MOTA	298	CB	ALA	60	9.762	0.122	-6.551	1.00 16.99
ATOM	299	N,	PHE	61	9.244	0.097	-9.571	1.00 13.66
ATOM	300	CA	PHE	61	9.423		-10.888	1.00 14.45
MOTA	301	C	PHE	61	9.413		-12.019	1.00 24.16
MOTA	302	0	PHE	61	9.881	_	-13.122	1.00 25.72
ATOH	303	CB	PHE	61	8.412		-11.119	1.00 15.76
MOTA	304	CG	PHE	61	8.907	-2.959	-10.671	1.00, 17.86
MOTA	305	CD1	PHE	61	8.887	-3.314	-9.320	1.00 19.64
HOTA	306	CD2	PHE	61	9.440	-3.854	-11.594	1.00 21.88
ATOM	307	CE1	PHE	61	9.396	-4.536	-8.905	1.00 22.58
MOTA	308	CE2	PHE	61	9.952	-5.080	-11.186	1.00 22.88
ATOM	309	CZ	PHE	61	9.929	-5.421	-9.842	1.00 21.09
ATOH	310	N	ALA	62	8.984	1.738	-11.699	1.00 20.90
MOTA	311	CA	ALA	62	8.961	2.830	-12.661	1.00 19.14
ATOM	312	С	ALA	62	10.359	3.437	-12.637	1.00 24.17
ATOM	313	0	ALA	62	10.842	3.949	-13.636	1.00 26.12
ATOH	314	СВ	ALA	62	7.936	3.865	-12.255	1.00 19.28
MOTA	315	N	MET	6,3	10.987	3.380	-11.472	1.00 18.82
MOTA	316	CA	MET	63	12.334	3.883	-11.258	1.00 17.35
MOTA	317	C	MET	63	13.304	2.892	-11.889	1.00 23.98
ATOM	318	0	MET	63	14.313	3.286	-12.463	1.00 27.31
ATOM	319	СВ	MET	63	12.588	3.986	-9.753	1.00 19.12
ATOM	320	CG.	MET	63	14.018	4.202	-9.335	1.00 21.76
ATOM	321	SD	MET	63 ⁻	14.350	3.308	-7.827	1.00 25.88
ATOM	322	CE	MET	63	14.496	1.636	-8.499	1.00 22.73
MOTA	323	N	LEU	64	12.946	1.614	-11.842	1.00 18.80
ATOM	324	CA	LEU	64	13.771	0.544	-12.398	1.00 18.07
MOTA	325	С	LEU	64	13.683	0.510	-13.903	1.00 23.00
MOTA	326	0	LEU	64	14.634	0.119	-14.573	1.00 24.06
MOTA	327	ĊB	LEU	64	13.328	-0.814	-11.853	1.00 17.89
MOTA	328	CG	LEU	64	14.035	-2.059	-12.384	1.00 22.43
ATOM	329	CD1	LEU	64	15.469	-2.072	-11.924	1.00 22.35
ATOM	330	CD2	LEU	64	13.307	-3.307	-11.918	1.00 25.46
MOTA	331	N	SER	65	12.551	0.949	-14.437	1.00 20.59
MOTA	332	CA	SER	. 65	12.361	0.942	-15.874	1.00 20.79
MOTA	333	C ·	SER	65	13.360	1.880	-16.533	1.00 24.97
MOTA	334	0	SER	65	13.646	1.751	-17.716	1.00 26.49
MOTA	335	CB	SER	65	10.947	1.367	-16.230	1.00 24.46
MOTA	336	Œ	SER	65	10.758	2.745	-15.960	1.00 37.12
MOTA	337	N	LEU	66	13.887	2.826	-15.763	1.00 18.20
MOTA	338	CA	LEU	66	14.848	3.781	-16.292	1.00 16.09
ATOM	339	С	LEU	66	16.145		-16.658	1.00 19.61
MOTA	340	0	LEU	66	16.942		-17.409	1.00 19.59
MOTA	341	CB	LEU	66	15.130	4.892	-15.274	1.00 15.19
ATOM	342	CG	LEU	66	13.921	5.676	-14.767	1.00 17.85
ATOM	343	CD1	LEU	66	14.335	6.635	-13.683	1.00 17.63
MOTA	344	CD2		66	13.251	6.396	-15.902	1.00 16.61
ATOM	345	N	GLY	67	16.370	1.904	-16.096	1.00 18.62
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ATOM	346	CA	GLY	67	17.582	1.159 -16.384	1.00 19.69
ATOM	347	С	GLY	67	17.302	0.034 -17.361	1.00 25.74
ATOM	.348	0	GLY	67	18.188	0.756 -17.691	1.00 27.68
ATOM	349	N	THR	- 68	16.063	-0.038 -17.832	1.00 18.44
MOTA	350	CX	THR	68	15.677	-1.079 -18.769	1.00 15.20
ATOM	351	С	THR	~ 68	15.511	-0.525 -20.177	1.00 16.21
ATOM	352	0	THR	68	15.277	0.671 -20.357	1.00 16.20
MOTA	353	CB	THR	68	1,4.367	-1.777 -18.321	1.00 10.45
MOTA	354	OG1	THR	68	13.305	-0.815 -18.219	1.00 9.49
ATOM	355	CG2	THR	68	14.557	-2.454 -16.987	1.00 1.57
ATOM	356	N	LYS	69	15.575	-1.408 -21.165	1.00 11.62
ATOM	357	CA	LYS	69	15.435	-1.009 -22.557	
ATOM	358	С.	LYS	69	14.490	-1.962 -23.291	1.00 20.44
ATOM	359	0	LYS	69	14.153	-3.029 -22.762	1.00 21.50
ATOM	360	CB	LYS	69	16.815	-1.000 -23.238	1.00 14.74
ATOM	361	CG	LYS	69	17.764	0.048 -22.676	1.00 30.38
ATOM	362	CD	LYS	69	19.066	0.138 -23.445	1.00 43.51
ATOM	363	CE	LYS	69	19.975	1.219 -22.860	1.00 64.68
ATOM	364	NZ	LYS	69	21.244	1.376 -23.619	1.00 74.82
ATOM	365	N	ALA	-70	13.948	-1.521 -24.426	1.00 16,85
	366	CA	ALA	70	13.083	-2.372 -25.248	1.00 17.01
MOTA	367	C	ALA	70	11.969	-3.131 -24.526	
ATOM	368	٥	ALA	70	11.415	-2.636 -23.551	1.00 23.28
ATOM	369	СВ	ALA	.70	13.933	-3.341 -26.054	1.00 18.23
ATOM	370	N	ASP	71	11.653	-4.333 -25.010	1.00 18.01
ATOM	371	CA	ASP	71	10.585	-5.172 -24.451	1.00 18.92
ATOM	372	С	ASP	.71	10.616	-5.216 -22.920	1.00 26.30
MOTA	373	0	ASP	71	9.603	-4.985 -22.263	1.00 27.15
MOTA	374	СВ	ASP	71	10.672	-6.612 -24.988	1.00 22.13
ATOM	375	CG	ASP	71	10.770	-6.686 -26.513	1.00 51.31
ATOM	376		ASP	71	10.518	-5.694 -27.221	1.00 59.09
ATOM	377		ASP	71	11.099	-7.783 -27.013	1.00 60.06
ATOM	378	N	THR	. 72	11.812	-5.438 -22.371	1.00 24.21
ATOM	379	CA	THR	72	12.044	-5.535 -20.925	1.00 22.94
ATOM	380	С	THR	72	11.427	-4.353 -20.167	1.00 24.66
ATOM	381	0	THR	72	10.811	-4.517 -19.112	1.00 24.36
ATOM	382	СВ	THR	72	13.569	-5.555 -20.653	1.00 29.44
ATOM	383	0G1	THR	72 .	14.173	-6.639 -21.376	1.00 30.43
MOTA	384	CG2	THR	72	13.862	-5.721 -19.167	1.00 28.97
ATOM	385	N	HIS	73	11.600	-3.172 -20.743	1.00 19.21
ATOM	386	CA	HIS	73	11.120	-1.931 -20.187	1.00 17.25
MOTA	387	С	HIS	73	9.629	-1:737 -20.394	1.00 24.90
ATOM	388	Ο.	HIS	.73	8.888	-1.507 -19.441	1.00 25.89
ATOM	389	CB	HIS	73	11.892	-0.798 -20.848	1.00 16.14
ATOM.	390	CG	HIS	73	11.303	0.545 -20.624	1.00 18.99
MOTA	391	ND1	HIS	73	10.438	1.137 -21.525	1.00 21.51
ATOM	392		HIS	73	11.467	1.437 -19.622	1.00 21.38
ATOM	393		HIS	73	10.103	2.334 -21.083	1.00 21.60
ATOM	394		HIS	73	10.713	2.539 -19.936	1.00 21.99
ATOM	395	N	ASP	74	9.209	-1.800 -21.656	1.00 21.47
ATOM	396	CA	ASP	74	7.814	-1.609 -22.053	1.00 19.79
MOTA	397	С	ASP	74	6.922	-2.530 -21.279	1.00 21.39
ATOM	398	0	ASP	74	5.805	-2.167 -20.931	1.00 23.80
ATOM	399	СВ	ASP	74	7.650	-1.873 -23.540	1.00 21.48
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ATOM	400 CG	ASP 74	8.272 -0.790 -24.394 1.00 31.57
MOTA	401 OD1	ASP 74	8.786 0.212 -23.867 1.00 31.06
MOTA	402 OD2	ASP 74	8.230 -0.945 -25.629 1.00 42.19
ATOM	403 N	GLU 75	7.455 -3.702 -20.973 1-00 14.48
MOTA		GLU 75	6.738 -4.701 -20.221 1.00 14.26
MOTA	405 C	GLU 75	6.482 -4.248 -18.788 1.00 20.06
MOTA		GLU 75	5.425 -4.543 -18.226 1.00 21.61
ATOM		GLU 75	7.539 -5.989 -20.228 1.00 15.78
ATOM		GLU 75	6.812 -7.174 -19.656 1.00 32.90
ATOM		GLU 75	7.666 -8.413 -19.711 1.00 59.73
ATOM	410 OE1 (8.603 -8.484 -18.886 1.00 56.25
ATOM	411 OE2 (7.419 -9.301 -20.548 1.00 53.12
ATOM		ILE 76	7.445 -3.542 -18.196 1.00 15.73
MOTA		LE 76	7.290 -3.055 -16.828 1.00 15.03
MOTA		LE 76	6.214 -1.984 -16.793 1.00 21.66
MOTA MOTA		LE 76	5.222 -2.122 -16.079 1.00 22.85
ATOM	_	LE 76	8.614 -2.476 -16.251 1.00 17.25
ATOM	417 CG1 I		9.663 -3.582 -16.092 1.00 16.04
ATOM	418 CG2 I 419 CD1 I		8.362 -1.794 -14.914 1.00 17.21
ATOM	,		10.958 -3.133 -15.411 1.00 14.11
ATOM		EU 77 EU 77	6.382 -0.941 -17.599 1.00 16.68
ATOM		EU 77	5.406 0.148 -17.633 1.00 15.81
ATOM		EU 77	4.007 -0.357 -17.981 1.00 19.88
ATOM	_	EU 77	3.017
ATOM		רל טב דל טב	
ATOM	426 CD1 L		1.00 19.72
MOTA	427 CD2 LI		
ATOM	428 N G	· ·	7.199 2.525 -16.901 1.00 17.87 3.930 -1.264 -18.942 1.00 15.10
ATOM	429 CA GI		2.646 -1.819 -19.325 1.00 14.48
MOTA	430 C GI		2.115 -2.691 -18.194 1.00 15.59
MOTA	431 0 GI		0.908 -2.773 -17.979 1.00 13.66
ATOM	432 CB GL		2.767 -2.619 -20.608 1.00 16.58
MOTA	433 CG GL	บ 78	2.803 -1.805 -21.888 1.00 34.30
ATOM	434 CD GL		3.049 -2.682 -23.092 1.00 66.97
MOTA	435 OE1 GL		2.677 -3.877 -23.057 1.00 63.59
ATOM	436 OE2 GL		3.629 -2.174 -24.070 1.00 69.86
MOTA	437 N GL		3.027 -3.330 -17.471 1.00 14.27
ATOM	438 CA GL		2.649 -4.167 -16.341 1.00 16.07
MOTA MOTA	439 C GL		2.136 -3.316 -15.189 1.00 25.07
	440 0 GL		1.340 -3.768 -14.357 1:00 27.66
atom Atom	441 N LET		2.589 -2.066 -15.140 1.00 18.21
MOTA			2.143 -1.135 -14.108 1.00 14.93
ATOM			0.880 -0.410 -14.555 1.00 13.69
ATOM	444 0 LET 445 CB LET		0.553 0.668 -14.054 1.00 9.47
ATOM	446 CG LEU		3.227 -0.115 -13.769 1.00 14.43
ATOM	447 CD1 LEU		4.536 -0.689 -13.241 1.00 17.68
ATOM	448 CD2 LEU		5.522 0.440 -13.082 1.00 17.74
ATOM	449 N ASN		4.325 -1.444 -11.948 1.00 14.60 0.160 -1.027 -15.481 1.00 13.09
ATOM	450 CA ASN		
ATOM .	451 C ASN	81	
ATOM	452 0 ASN	81	
MOTA	453 CB ASN	81	
	-		-2.106 -0.407 -14.846 1.00 17.98

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ATOH	454	CG ASN	81	-2.96	8 -1.649 -14.76	1.00 44.40
MOTA	455	OD1 ASN		-4.06		
ATOM	456	ND2 ASN		-2.50		
ATOM	457	N. PHE	82.	0.05		
ATOM	458	CA PHE	82	0.342		
ATOM	459	C PHE	82	0.27		
ATOM	460	O PHE	. 82	1.019		
MOTA	461	CB PHE	82	1.729		
ATOM	462	CG -PHE	82	1.782		
MOTA	463	CD1 PHE	82	1.321		
ATOM		CD2 PHE	82	2.292		
MOTA		CE1 PHE	82	1.364		
MOTA		CE2 PHE	82	2.339		
ATOM		CZ PHE	82	1.874		
MOTA		n asn	83	-0.634	2.502 -20.542	1.00 16.94
ATOM		ca asn	83 .	-0.785	2.291 -21.972	1.00 15.03
ATOM		C ASN.	83	0.244	3.141 -22.670	1.00 22.28
MOTA		o asn	83	-0.001	4.310 -22.979	1.00 24.99
ATOM		CB ASN	83	-2.179	2.689 -22.429	1.00 16.41
ATOM		CG ASN	83	-2.427	2.344 -23.868	1.00 38.32
ATOM		DD1 ASN	83	-1.513	1.946 -24.587	1.00 32.12
ATOM		ND2 ASN	83	-3.668		1.00 35.98
ATOM ATOM		LEU	84	1.376	2.528 -22.978	1.00 17.89
		À LEU	84	2.474	3.248 -23.603	1.00 16.80
MOTA MOTA	478 C		84	2.171	3.967 -24.911	1.00 20.88
ATOM			84	2.840	4.945 -25.247	1.00 20.85
ATOM		B LEU G LEU	84	3.679	2.324 -23.764	1.00 15.58
ATOM		D1 LEU	84	4.156	1.723 -22.446	1.00 17.53
ATOM		D2 LEU	84 84	5.318	0.790 -22.689	1.00 16.86
ATOM	484 N		85	4.522	2.829 -21.484	1.00 15.95
ATOM	485 C		85	1.136 0.769	3.523 -25.610	1.00 18.16
ATOM	486 C		85	-0.057	4.122 -26.884 5.385 -26.715	
MOTA	487 0		85	-0.248		1.00 22.48
ATOM	488 C		85	-0.018	3.131 -27.758	1.00 22.41
ATOM		G1 THR	85	-1.145		1.00 39.97
ATOM		32 THR	85	0.874	1.978 -28.176	1.00 45.76 1.00 40.70
ATOM	491 N		86	-0.544	5.613 -25.507	1.00 18.29
MOTA	492 C		86	-1.367		1.00 17.94
MOTA	493 C	GLU	86	-0.731		1.00 17.34
ATOM	494 0	GLU	86	-1.012	8.991 -24.480	
MOTA	495 CE	GLU	. 86	-2.721		1.00 20.11
MOTA	496 CG	GLU	86	-3.508	5.480 -25.689	
MOTA	497 CD		86	-4.902		1.00 72.19
MOTA		1 GLU	86	-5.297		1.00 63.51
MOTA		2 GLU	86	-5598		1.00 73.21
NTOM	500 ห	ILE	87	0.152		1.00 15.47
NTOM	501 CA		87	0.806		1.00 16.06
MOTA	502 C	ILE	87	2.312		.00 21.27
TOM	503 0	ILE	87	2.974		00 21.63
MOT	504 CB		87	0.446		.00 19.45
MOT		1 ILE	87	1.034	9.101, -20.149 1	.00 19.82
MOT		2 ILE	87	0.925	6.624 -20.616 1	.00 19.35
TOM	507 CD:	1 ILE	87.	0.602		.00 34.75
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MOTA	508	3 N	PRO	88	2.86	8 9.508	-22.968	1.00 16	1 5
ATOM	509	CA	PRO	88	4.31		-23.152		
ATOM	510) C	PRO	88	- 5.02		-21.886		
ATOM	511	. 0	PRO	88	4.67		-20.783		
MOTA	512		PRO	88	4.52		-23.440		
MOTA	513	CG	PRO	8.8	3.19		-23.254		
MOTA	514	CD	PRO	88	2.14		-23.333	1.00 16.	
ATOM	515		GLU	89	6.03		-22.050	1.00 16.	
ATOM	516		GLU	89	6.78		-20.908	1.00 16.	
MOTA	517	_	GLU	89	7.33		-20.063	1.00 22.	
ATOM	518		GLU	89	7.49	7 8.823	-18.842	1.00 22.	
ATOM	519		GLU	89	7.89		-21.365	1.00 17.	
ATOM.	520		GLU	89	7.40		-22.033	1.00 34.	85
MOTA	521	CD	GLU	89	8.53		-22.483	1.00 61.	89
ATOH	522		GLU	89	9.710		-22.159	1.00 48.	17
ATOM	523		GLU	89	8.262			1.00 62.	
MOTA	524	N .	λLλ	90	7.636		-20.731	1.00 20.8	
ATOM	525	CA	ALA	90	8.156		-20.062	1.00 20.2	
MOTA	526	С	ALA	90	7.146		-19.016	1.00 22.0	
ATOM	527 528	0	ALA	90	7.525			1.00 19.2	
ATOM	529	CB N	ALA	90	8.392			1.00 21.1	
ATOM	530	CA.	GLN GLN	91 91	5.864			1.00 19.3	
ATOM	531	C	GLN	91	4.773			1.00 19.3	
ATOM	532	0	GLN	91				1.00 23.7	
ATOM	533		GLN	91	4.231 3.452	11.239		1.00 24.8	
ATOM	534		GLN	91	3.195	12.006		1.00 20.9	
ATOM	535		GLN	91	1.749	13.409 -		1.00 48.5	
ATOM	536	OE1		91	0.907	13.610 - 12.724 -		1.00 73.8	
ATOM	537	NE2		91	1.455	14.776 -		1.00 69.2	
ATOM	538		ILE		4.932		17.578	1.00 63.2	
ATOM	539		ILE	92	4.823	8.557 -		1.00 17.49 1.00 15.82	
ATOM	540		ILE	92	5.773	8.855 -		1.00 21.60	
ATOM	541		ILE	92	5.363	8.946 -		1.00 21.60	
ATOM	542		ILE	92	5.160	7.164 -		1.00 22.63	
ATOM	543		ILE	92	4.097	6.754 -		1.00 17.24	
ATOM	544	CG2	ILE .	92	5.295	6.150 -		1.00 16.93	
ATOM	545	CD1	ILE	92	4.330	5.414 -		1.00 23.97	
MOTA	546	N F	HIS	93	7.035	9.082 -		1.00 17.87	
ATOM	547	CA F	iis	93	8.055	9.374 -		00 16.61	
MOTA	548	CF	iis	93	7.815	10.736 -		.00 21.94	
ATOM	549	0 F	IIS	93	7.971	10.877 -		.00 21.04	
ATOM	550		IIS	93	9.449	9.289 -1		.00 16.45	
MOTA			IIS	93	9.768	7.933 -1	'	.00 19.48	
MOTA		ND1 H		93	9.691	6.775 -1		.00 21.37	
MOTA		CD2 H		93	10.138	7.557 -1		.00 21.86	-
MOTA		CE1 H		93	9.994	5.737 -1	5.962 1	.00 21.40	
ATOM		NE2 H		93	10.271	6.186 -1	7.177 1	.00 22.02	
MOTA			LU	94	7.382	11.720 -1	4.901 1	.00 21.50	
MOT			LU	94	7.111	13.062 -1	4.370 1	.00 21.71	
MOT			LU	94	5.996	12.961 -1	3.338 1	.00 27.52	
TOM			LU	94	6.059	13.576 -1		.00 29.26	
TOM			LU	94		14.023 -1	5.488 1	.00 23.76	
TOM	561	CG G	LU .	94	6.268	15.417 -1	5.028 1	.00 46.66	

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ATON	562	CE	GLU	94		5.732	16.286	-16.166	1.00100.99
MOTA	563	OE	1 GLU	94		5.513	15.769	-17.285	
MOTA	564	. · OE	2 GLU	94		5.517	17.493	-15.933	1.00109.77
MOTA	565	N	GLY	95		5.015	12.116	-13.632	1.00 22.71
MOTA	566	CA	GLY	95		. 3.903	11.932	-12.723	1.00 21.91
MOTA	567	C	GLY	95		4.352	11.312	-11.407	1.00 26.12
ATOM	568	0	- GLY	95		3.928	11.757	-10.337	1.00 26.39
ATOM	569	N	PHE	96		5.247	10.324	-11.477	1.00 20.03
ATOM	570	CX	PHE	96		5.755	9.640	-10.283	1.00 17.73
MOTA	571	C	PHE	96		6.559	10.556	-9.384	1.00 23.93
ATOM	572	0	PHE	96		6.513	10.436		
ATOM	573	CB	PHE	96		6.598	8.413	-10.656	
MOTA	574	CG	PHE	96		5.784		-10.895	
MOTA	575		1 PHE	96		5.127			1.00 23.98
MOTA	576	CD	2 PHE			5.627		-12.172	1.00 19.50
MOTA	577	CE	1 PHE	96		4.323		-10.057	1.00 24.57
ATOM	578	CE		96		4.825		-12.397	
ATOM	579	CZ	PHE	96	•	4.174		-11.339	1.00 21.77
MOTA	580	N	GLN	. 97		7.273	11.493	-9.994	1.00 20.80
MOTA	581	CA	GLN	97		8.085	12.438	-9.254	1.00 19.78
MOTA	582	С	GLN	97		7.180	13.352	-8.453	1.00 21.06
ATOM	583	0	GLN	97		7.410	13.575	-7.267	1.00 19.65
ATOM	584	CB	GLN	97		8.961		-10.212	
ATOM	. 585	CG	GLN	97		10.039		-10.901	1.00 53.46
MOTA	586,	CD	GLN	97	•	10.845	13.187		1.00 97.50
MOTA	587	CE		97		10.669		-12.070	1.00102.38
ATOM	588		GLN	97		11.729		-12.616	1.00 92.80
ATOM	589	N	GLU	98		6.099	13.804	-9.078	1.00 18.84
ATOM	590	CA	GLU	98		5.156	14.693	-8.398	1.00 20.12
MOTA	591	C	GLU	98		4.676	14.065	-7.090	1.00 27.71
ATOM	592	0	GLU	98		4.631	14.715	-6.035	1.00 29.41
ATOM	593	CB	GLU	98		3.927	14.991	-9.276	1.00 21.69
ATOM	594	CG	GLU	98		4.158		-10.419	1.00 43.54
ATOM	595 596	CD	GLU	. 98		4.424	17.394	-9.964	1.00 92.78
ATOM ATOM	596 597		GLU	98		4.622	17.646 18.275	-8.753	1.00 95:84
ATOM	598	N N	GLU	98 99		4.448 4.384	12.775	-7.171	1.00103.56
ATOM	599	CA	LEU	99		3.893			1.00 21.26
ATOM	600		LEU	99		4.910	11.999	-6.053	1.00 19.05
ATOM	601	0	LEU LEU	99		4.587	11.927 12.186	-4.934 -3.771	1.00 24.77
ATOM	602	CB	LEU	99		3.552	10.599	-5.771 -6.541	1.00 24.94 1.00 18.47
ATOM	603	CG		99		2.828	9.714		
ATOM	604		LEU LEU	99		1.698	10.478	-5.548 -4.922	1.00 23.63 1.00 23.11
ATOM .	-605		LEU	99		2.360	8.461	-6.253	1.00 23.11
ATOM	506	N N	LEU	100		6.155	11.657	-5.303	1.00 27.37
ATOM	607	CA	LEU	100		7.240	11.541	-4.331	1.00 20.15
ATOM	608	C	LEU	100			.12.896	-3.703	1.00 20.15
ATOM	609	0	LEU	100		7.939	12.970	-2.540	1.00 23.86
MOTA	610	CB	LEU.	100		8.491	10.962	-5.006	1.00 20.24
ATOM	611	cc	LEU	100		8.299	9.633	-5.750	1.00 25.35
ATOM	612		LEU	100		9.589	9.234	-6.433	1.00 26.21
ATOM	613		LEU	100		7.823	8.558	-4.797	1.00 26.62
ATOM	614	N	ARG	101	-	7.259	13.960	-4.453	1.00 24.56
ATOM	615	CA	ARG	101		7.494	15.338	-4.019	1.00 26.50
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HOTA	616	С	ARG	101	6.535	15.594	-2.869	1.00 29.95
MOTA	617	0	ARG		6.894	16.189	-1.840	1.00 29.61
ATOH	618	CB	ARG	101	7.196	16.303	-5.172	1.00 31.10
ATOM	619	CG	ARG	101	7.540	17.759	-4.904	1.00 50.33
ATOM	620	CD	ARG	101	6.916	18.678	-5.963	. 1.00 78.18
ATOM	621	NE	ARG	101	6.993	20.088	-5.582	1.00 97,33
ATOM	622		ARG	101	6.475	20.588	-4.462	1.00110.00
MOTA	623	NH	1 ARG	101	`5.838	19.798	-3.606	
MOTA	624	NH.	2 ARG	101	6.619	21.876	-4.183	
ATOM	625	N	THR	102	5.322	15.087	-3.046	1.00 26.75
ATOH	626	CA	THR	102	4.297	15.221	-2.042	1.00 27.11
MOTA	627	С	THR	102	4.624	14.273	-0.900	
MOTA	628	0	THR	102	4.400	14.585	0.236	1.00 34.20
MOTA	629	CB	THR	102	2.907	14.861	-2.598	1.00 46.57
ATOM	630	OG:		102	2.619	15.665	-3.757	1.00 50.68
ATOM	631	CG		102	1.841	15.098	-1.543	
HOTA	632	N	LEU	103	5.202	13.122	-1.176	1.00 24.28
MOTA	633	CA	LEU	103	5.490	12.223	-0.080	1.00 21.81
MOTA	634	C:	ĻEU	103	6.731	12.644	0.744	1.00 28.18
MOTA	635	0	LEU	103	6.821	12.344	1.943	1.00 27.87
MOTA	636	CB	LEU	103	5.573	10.784	-0.605	1.00 20.79
MOTA	637	CG	LEU	103	4.304	10.284	-1.322	1.00 24.82
ATOM	638		LEU	103	4.471	8.845	-1.787	1.00 25.29
ATOM	639		LEU	103	3.104	10.398	-0.409	1.00 25.55
ATOM	640	N	ASN	104	7.657	13.385	0.127	1.00 26.19
MOTA	641 642	CA C	ASN	104	8.876	13.829	0.822	1.00 26.92
ATOM	643	0	ASN	104	8.494	14.854	1.879 3.077	1.00 29.91
ATOM	644		asn Asn	104	8.663 9.890	14.622 14.438	-0.156	1.00 30.79 1.00 33.86
ATOM	645	CG	ASN	104 104	11.321	13.946	0.087	1.00 33.86
ATOM	646		ASN	104	12.120	13.843	-0.853	1.00 64.96
ATOM	647		ASN	104	11.647	13.632	1.341	1.00 57.77
ATOM	648	N	GLN	105	7.999	15.994	1.419	1.00 25.50
ATOM	649	CA	GLN	105	7.532	17.062	2.315	1.00 24.94
ATOM	650	С	GLN	105	6.172	17.475	1.746	1.00 28.97
ATOM	651	0	GLN	105	6.061	18.452	0.978	1.00 31.34
ATOM	652	CB.	GLN	105	8.540	18.227	2.379	1.00 26.65
MOTA	653	CG	GLN	105	8.082	19.454	3.220	1.00 66.64
ATOM	654	CD	GLN	105	7.531	19.104	4.601	1.00105.25
ATOM	655	OE1	GLN	105	7.741	18.003	5.110	1.00107.04
ATOM	656	NE2	GLN	105	6.794	20.042	5.200	1.00 99.00
MOTA	657	N	PRO	106	5.125	16.697	2.076	1.00 20.86
ATOM	658	CA	PRO	106	3.741	16.897	1.634	1.00 18.74
MOTA	659	С.	PRO	106	3.006	18.152	2.061	1.00 22.19
MOTA	660	. 0	PRO.	106	3.582	19.222	2.292	1.00 21.82
MOTA	661	CB	PRO	106	3.007	15.630	2.128	1.00 19.72
HOTA	662	CG	PRO	106		15.067	3.172	1.00 24.39
MOTA	663	CD	PRO	106		15.488	2.912	1.00 20.20
ATOM .	664	N	ASP	107		17.981	2.058	1.00 18.09
MOTA	665	CA	ASP	107		18.968	2.471	1.00 17.45
ATOM	666	C	ASP	107		18.789	3.996	1.00 21.97
ATOM	667	0 ,	ASP	107		19.441	4.670	1.00 21.75
ATOM	668	CB	ASP	107		18.596	1.818	1.00 18.93
ATOM	669	CC	ASP	107	-1.728	19.659	1.968	1.00 27.12

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ATOM	670	OD	1 ASP	107		-1.630	20.613	2.757	1.00	27.68
ATOM	671	OD:	2 ASP	107		-2.746	19.503	1.264	1.00	32.41
MOTA	672	N	SER	108		1.516	17.945	4.522	1.00	18.59
ATOM	673	CA	SER	108		1.615	17.640	5.945	1.00	18.81
MOTA	674	C	SER			2.474	16.396	6.169		24.34
MOTA	675	. 0	SER	108		2.385	.15.429	5.416		23.92
ATOM	676	CB	SER			0.225	17.412	6.549		22.39
MOTA	677	œ	SER	108		0.291	17.068		1.00	
ATOM	678	N	GLN	109		3.269		7.230		22.70
ATOM	679	CA	GLN	109		4.127	15.294	7.567		22.48
ATOM	680	С	GLN			3.261	14.181	8.176		28.45
ATOM	681	0	GLN	109		2.765	14.282	9.305		29.12
ATOM	682	CB	GLN	109		5.228	15.755	8.528		23.92
MOTA	683	CG	GLN	109		6.103	14.639	9.058		60.78
MOTA	684	CD	GLN			7.177	15.138	10.006		02.33
ATOM	685		GLN	109		7.171	16.298	10.434		99.17
MOTA	686 687	NE2	GLN LEU	109		8.106	14.257	10.346		23.12
ATOM	688	N CA		110		3.019 2.253	13.171	7.702		21.02
ATOM	689	C.	LEU LEU	110 110		3.304	11.985 10.897	7.752		26.33
ATOM	690	0	LEU	110		4.126	10.628	7.070		25.47
ATOM	691	СВ	LEU	110	-	1.397	11.615	6.494	1.00	
ATOM	692	CG	LEU	110		0.740	10.249	6.412	1.00	
ATOM	693		LEU	110		-0.593	10.264	7.141	1.00	
ATOM	694		LEU	110		0.542	9.904	4.949	1.00	
ATOM	695	N	GLN	111		3.329	10.300	9.143	1.00	
ATOM	696	CA	GLN	111		4.340	9.272	9.397	1.00	
ATOM	697	Ċ	GLN	111		4.256	8.154	8.383	1.00	30.55
ATOM	698	0	GLN	. 111		3.314 -	7.360	8.388	1.00	
MOTA	699	CB	GLN	111		4.271	8.702	10.819	1.00	27.74
ATOM	700	CG	GLN	111		4.825	9.658	11.876	1.00	70.63
ATOM	701	CD	GLN	111	• •	5.296	8.965	13.140	1.001	07.33
ATOM	702		GLN	111	•	5.247	7.736	13.265	1.001	
ATOM	703		GLN	111		5.787	9.758	14.081	1.001	
MOTA	704	N	LEU	112		5.222	8.154	7.477	1.00	
ATOM	705	CA	LEU	112		5.289	7.152	6.441	1.00	
ATOM	706	С	LEU	112		6.746	6.771	6.330		27.64
MOTA	707	0	LEU	112		7.605	7.617	6.082		31.14
ATOM	708	CB	LEU	112		4.786	7.712	5.108		19.99
MOTA	709 710	CG	LEU	112		4.265	6.699	4.092 4.769		25.35
ATOM ATOM	711	CD1	LEU	112 112		3.255 3.631	5.811 7.405	2.930	1.00 2	
ATOM	712	N	LEU THR	113		7.022	5.500	6.569	1.00 2	
ATOM	713	·CA	THIR	113		B.377	4.992	6.497	1.00 2	
ATOM	714	C	THR	113		8.426	4.105	5.263	1.00 2	
ATOM	715	ō	THR	113		8.173	2.903	5.349	1.00 2	
ATOM	716	СВ	THR	113		8.693	4.173	7.744	1.00 3	
ATOM	717	0G1		.113		8.266	4.902	8.901	1.00 3	
ATOM	718	CG2		113	. 1	0.178	3.922	7.843		2.13
ATOM	719	N	THR	114		8.755	4.699	4.121	1.00 1	
ATOM	720	CA	THR	114		8.790	3.956	2.875	1.00 1	
MOTA	721	C	THR	114		0.154	4.090	2.247	1.00 2	
ATOM .	722	٥	THR	114		0.800	5.126	2.394	1.00 2	6.47
MOTA	723	CB.	THR .	114		7.757	4.503	1.897	1.00 3	

MOTA	724		THR		6.481				
MOTA	725		THR		7.663			1.00 2	
MOTA	726		GLY		10.598			_	
ATOM	727		GLY		11.891			1.00 1	
MOTA	728		GLY	115	12.062	2.062	-0.098	1.00 1	7.27
MOTA	729	0	GLY	115	11.266	1.133	-0.168	1.00 1	5.25
MOTA	730	N	ASN	116	13.091	2.201	-0.916	1.00 1	4.85
MOTA	731	CA	ASN	116	13.362	1.224	-1.938	1.00 1	5.66
MOTA	. 732	C	ASN	116	14.855	1.125	-1.986	1.00 2	2.82
ATOM	733	0	ASN	116	15.540	2.139	-2.127	1.00 2	4.97
MOTA	734	CB	asn	116	12.841	1.707	-3.288	1.00 1	5.07
ATOM	735	CG	asn	116	12.611	0.571	-4.262	1.00 3	2.48
ATOM	. 736	001	ASN	116	- 11.539	-0.044	-4.302	1.00 1	9.35
ATOM	737	ND2	ASN	116	13.635	0.269		1.00 20	5.27
ATOM	738	N	GLY	117	15.362	-0.071		1.00 1	
ATOM	739	CA	GLY	117	16.793	-0.279		1.00 1	
MOTA	740	С	GLY	117	17.191	-1.264		1.00 19	
ATOM	741	0	GLY	117	16.601	-2.334		1.00 19	
MOTA	742	N	LEU	118	18.142	-0.863		1.00 16	
ATOM	743	CA	LEU	118	18.639	-1.733		1.00 18	
ATOM	744	C:	LEU	118	19.993	-2.254	-4.269	1.00 23	
ATOM	745	0	LEU	118	20.739	-1.547	-3.596	1.00 24	
ATOM	746	CB	LEU	118	18.789	-0.962	-6.031	1.00 19	
ATOM		CG	LEU	118	17.502	-0.316	-6.544	1.00 28	
ATOM	748	-	LEU	118	17.780	0.426	-7.840	1.00 29	
ATOM	749		LEU	118	16.424	-1.371	-6.744	1.00 32	
ATOM	750	N	PHE	119	20.280		-4.593	1.00 17	
ATOM	751	CA	PHE	119	21.541	-4.126	-4.210	1.00 16	
ATOM	752	c	PHE	119	22.134	-4.817	-5.418	1.00 22	
ATOM	753	o	PHE	119	21.631	-5.857	-5.855	1.00 22	
ATOM	754	СВ	PHE	119	21.313	-5.145	-3.095	1.00 17	
ATOM	755	CG	PHE	119	20.615	-4.580	-1.895	1.00 19	
ATOM	756	CD1		119	19.225	-4.559	-1.835	1.00 22	
ATOM	757		PHE	119	21.343	-4.032	-0.838	1.00 21	
ATOM	758		PHE	119	18.562	-3.997	-0.742	1.00 23	
ATOM	759		PHE	119	20.690	-3.465	0.265	1.00 24	
ATOM	760	cz	PHE	119	19.295	-3.447	0.312	1.00 21	
ATOM	761	N	LEU	120	23.187	-4.228	-5.970	1.00 18	
ATOM	762	CA	LEU	120	23.844	-4.781	-7.143	1.00 18	
ATOM	763	c	LEU	120	25.235	-5.300	-6.786	1.00 22	
ATOM	764	0	LEU	120		-4.822		1.00 22	
ATOM	765	СВ	LEU	120	23.940	-3.719	-8.240	1.00 19	
ATOM	766	CG	LEU	120	22.655	-3.047	-8.732	1.00 24	
ATOM	767		LEU	120	22.103	-2.051	-7.721	1.00 26	
ATOM	768		LEU	120	22.103	-2.335		1.00 28	
MOTA	769	N N		121			-7.556	1.00 26	
ATOM	770	CA	SER	121	25.702 26.993	-6.272 -6.894		1.00 16	
			SER						
MOTA	771 772	C	SER	121	28.199	-5.956	-7.33 4	1.00 23	
MOTA		O CP	SER	121	28.300	-5.060	-8.172	1.00 25	
MOTA	773	CB	SER	121	. 27.198	-8.031	-8.324	1.00 19.	
ATOM	774	OG N	SER	121	28.428	-8.698	-8.114	1.00 30.	
MOTA	775	N	GLU	122	29.083	-6.124	-6.355	1.00 19.	
ATOM	776	CA	GLU	122	30.323	-5.351	-6.313	1.00 19.	
MOTA	777	С	GLU	122	31.114	-5.991	-7.452	1.00 22.	. 25

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ATOM	778	0	GLU -	122	31.417	-7.182	-7.415	1.00 22.67
ATOM	779	CB	GLU	122	31.073	-5.563	-4.991	
ATOM:	780	CG	GLU	122	30.907	-4.444	-3.958	1.00 39.74
MOTA	781	CD	GLU	122	31.566	-4.756	-2.618	1.00 71.75
MOTA	782	OEI	GLU	122	31.886	-5.935	-2.354	1.00 64.09
MOTA	783	OE2	GLU .	122	31.753.	-3.819	-1.818	1.00 68.63
MOTA	784	N	GLY	123	31.369	-5.223	-8.497	1.00 18.92
ATOM	785	CA	GLY	123	32.095	-5.750	-9.637	1.00 19.03
MOTA	786	C .	GLY	123	31.430	-5.342	-10.937	1.00 24.70
ATOM	787	0	GLY	123	32.028	-5.437		1.00 25.16
MOTA	788	N	LEU	124	30.169	-4.931	-10.859	1.00 19.85
ATOM	789	CY	LEU	124	29.457	-4.493	-12.048	1.00 18.58
MOTA	790	С	LEU	124	29.702	-2.997	-12.249	1.00 21.59
MOTA	791	0	LEU	124	29.545		-11.318	
MOTA	792	CB	LEU	124	27.950		-11.928	1.00 17.99
ATOM	793	CG	LEU	124	27.479	-6.233	-11.813	1.00 22.20
MOTA	794		LEU	124	25.964	-6.268		1.00 21.86
ATOM	795.	CD2	LEU	124	28.003	-7.073		1.00 25.97
ATOM	796	N	LYS	125	30.131	-2.625		1.00 18.27
ATOM	797	CA	LYS	125	30.384		-13.756	1.00 19.18
MOTA	798	C	LYS	125	29.060	-0.571		1.00 28.77
ATOM	799	0	LYS	125	28.658	-0.505		
ATOM	800	СВ	LYS	125	31.384	-1.081	-14.901	1.00 48.12
MOTA	801	CG	LYS	125	31.786	0.355	-15.201 -16.452	1.00 72.20
MOTA	802	CD	LYS	125	32.644		-16.452	1.00 72.20
MOTA	803	CE	LYS	125	32.959			1.00110.00
ATOM	804	NZ	LYS	125	31.717 28.342	2.597 -0.187		1.00 20.76
ATOM	805	N	LEU	126	28.342		-13.039	1.00 19.13
MOTA	806	CA	LEU	126	27.043		-13.191	1.00 20.04
ATOM	807	C	LEU	126 126	28.085		-13.858	1.00 19.63
ATOM	808	0	LEU	126	26.416		-11.808	1.00 18.62
MOTA	809 810	CB	LEU LEU	126	26.553		-10.794	1.00 22.87
MOTA MOTA	811		LEU	126	25.839	-0.077		1.00 23.68
ATOM	812		LEU	126	26.012		-11.344	1.00 24.83
ATOM	813	N	VAL	127	25.995	2.109		1.00 16.04
ATOM	814	CA	VAL	127	25.861	3.348	-15.356	1.00 16.40
ATOM	815	C	VAL	127	25.538	4.485	-14.392	1.00 22.64
ATOM	816	ō	VAL	127	24.529	4.463	-13.696	1.00 24.15
ATOM	817	СВ	VAL	127	24.759	3.223	-16.409	1.00 20.46
ATOM	818	CG1		127	24.518	4.554	-17.101	1.00 21.01
ATOM	819	CG2		127	25.139		-17.423	1.00 20.09
ATOM	820	N	ASP	128	26.391	5.497	-14.367	1.00 19.51
ATOM		CA	ASP -	128	26.211	6.629	-13.468	1.00 19.12
ATOM	822	Ċ	ASP	128	24.851	7.287	-13.620	1.00 23.48
MOTA	823	0	ASP	128	24.201		-12.630	1.00 24.08
MOTA	824	CB	ASP	128	27.329		-13.675	1.00 21.97
MOTA	825	CG	ASP	128	28.701		-13.525	1.00 44.91
MOTA	826	OD1		128	29.016		-12.448	1.00 47.60
MOTA	827	OD2	ASP	128	29.467		-14.519	1.00 53.48
MOTA	828	N	LYS	129	24.387		-14.859	1.00 19.51
MOTA	829	CA	LYS	129	23.096		-15.126	1.00 18.71
MOTA	830	C	LYS	129	21.985		-14.334	1.00 23.17
MOTA	831	0	LYS	129	21.214	8.051	-13.663	1.00 24.46

ATOH	83	2 C	B LYS	129		22.78	6 8.0	08 -16.6	25 1	00 20	79
MOTA	- 83	3 CC	LYS	129		21.45		43 -16.99		00 29	
ATOM	83	4 CI	LYS	129.		21.32		74 -16.49		00 29	
ATOM	83	S CE	LYS	129		19.94		22 -16.78		00 32	
MOTA	83	6 N2	LYS	129		19.80		2 -16.43	_	00 32	
ATOM	83	7 N	PHE	130		21.94		18 -14.37		00 18	
ATOM	. 83	B. CA	PHE	130		20.91		6 -13.64		00 17	
MOTA	839	C	PHE	130		20.98		3 -12.16		00 24	
ATOM	840	0	PHE	130		19.96		1 -11.54		00 26	
MOTA	841	CB	PHE	130		21.04		9 -13.90		00 17	
MOTA	842			130		20.05		8 -13.13		00 18	
MOTA	843		1 PHE	130		18.689	3.10	5 -13.38		00 21	
ATOM	844		2 PHE	130		20.480	2.09	8 -12.15		00 20	
MOTA	845		1 PHE	130		17.775	2.35	5 -12.67		00 21	
ATOM	846		2 PHE	130		19.574	1.34	4 -11.43		00 23	
MOTA	847		PHE	130		18.218	1.47	1 -11.70		0 22	
MOTA	848		LEU	131		22.198		3 -11.60°	7 1.0	0 19	
MOTA	849		LEU	131		22.390	5.86	9 -10.18		0 18	
ATOM	850	-	LEU	131		21.960		2 -9.85	1.0	0 24	. 96
ATOM	851	0		131		21.547			1.0	0 25	
MOTA	852	CB	LEU	131		23.840				0 18.	. 23
ATOM	853	CG	LEU ·			24.277		3 -10.024		0 23.	
ATOM	854		LEU	131		25.721		-9.608		0 23.	
MOTA	855		LEU	131		23.369				0 24.	
ATOM	856	N	GLU	132		22.032		-10.851		0 21,	
ATOM	857	CA	GLU	132		21.603		-10.706		0 20.	
ATOM	858	C	GLU	132		20.089		-10.532		0 23.	
ATOM	859	0	GLU	132		9.558	10.012			0 23.	
ATOM ATOM	860 861	CB	GLU	132		1.938		-11.964		0 22.	
ATOM	862	CG	GLU	132		3.416		-12.279		39.	
ATOM	863	CD	GLU	132		3.628		-13.548		72.	
ATOM	864	OE2	GLU .	132		2.636		-14.100		55.	
ATOM	865	N N	ASP	132 133		4.788		-13.995		75.	
ATOM	866	CA	ASP	133		9.420 7.958		-11.510		19.	
ATOM	867	C	ASP	133		7.394		-11.556		19.	
ATOM	868	ō	ASP	133		6.344	8.517	-10.303		23.5	
ATOM	869	СВ	ASP	133		7.543		-9.793 -12.793		25.1	
ATOM	870	CG	ASP	133		7.978		-14.102		22.7	
ATOM	871	OD1		133		8.560		-14.102		34.2	
ATOM	872	OD2		133		7.719		-15.158		34.7	
ATOM	873	N	VAL	134	_	B.110	7.126	-9.806		38.4	
ATOM	874	CA	VAL	134		7.694	6.420	-8.605		17.7 17.4	
MOTA	875	С	VAL	134		7.727	7.372	-7.397		24.7	
ATOM -	876		VAL	134		5.676	7.667	-6.829		24.5	
MOTA	877		VAL	134		3.576	5.171	-8.358		19.7	
MOTA	878	CG1		134		3.265	4.560	-7.010		18.8	
MOTA	879	CG2		134		3.343	4.151	-9.454		19.6	
MOTA	880		LYS	135		.902	7.940	-7.106		22.5	
NOTA	881		LYS	135		.086	8.846	-5.965		22.4	
NTOM	882		LYS	135		.350	10.174	-6.089		26.4	
NOT	883		LYS	135		.578	10.556	-5.201		27.6	
MOTA	884		LYS	135		.573	9.123	-5.734	1.00		
MOTA	885	CG :	LYS	135		. 253	8.129	-4.805	1.00		
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MOTA	886	CD	LYS	135	21.299	6.711	_	1.00	45.73
MOTA	887			135	21.886	5.750			58.07
MOTA	888			135	21.078	5.726		1.00	74.45
ATOM	889		LYS	136	18.669	10.916			21.97
ATOM	890			136	18.044	12.207			22.48
ATOM	891		LYS	136	. 16.573	12.089			26.04
MOTA	892		LYS	136	15.709	12.328			27.39
ATOM	893			136	18.784	12.981			27.45
ATOM	894			136	20.077	13.637			57.41
MOTA	895			136	20.843	14.190			70.91
MOTA	896	CE	LYS	136	21.349		-10.055		78.67
MOTA	897	NZ	LYS	136	21.862		-11.389		83.40
MOTA	898	N	LEU	137	16.289	11.710			19.69
MOTA	899	CX	LEU		14.911				18.86
ATOM	900	C	LEU	137	13.965	10.854			25.85
ATOM	901	0	LEU	137	13.037	11.462	-7.987		31.02
ATOH	902	CB	LEU	137	14.837		-10.842		18.94
ATOM	903	CG	LEU	137	15.392		-12.026		25.53
MOTA	904		LEU	137	15.075		-13.276		26.83
MOTA	905		LEU	137	14.799		-12.098		28.13
ATOM	906	N	TYR	138	14.227	9.575	-8.250		18.00
ATOM	907	CA	TYR	138	13.340	8.793	-7.405		15.91
ATOM	908	С	TYR		13.736	8.609	-5.950		19.32
ATOM	909	0	TYR	138	13.063	7.879	-5.225		16.98
MOTA	910	CB	TYR	138	13.045	7.441	-8.052		16.71
ATOM	911	CG	TYR	138	12.114	7.486	-9.256		19.54
MOTA	912	CD1		138	12.488		-10.449	1.00	
ATOM	913		TYR	138	10.863	6.872	-9.213	1.00	
ATOM ATOM	914 915	CEI		138	11.631		-11.564	1.00	
MOTA	916	CE2		138 138	10.003		-10.324 -11.489	1.00	
MOTA	917	OH	TYR TYR	138	9.550		-12.570	1.00	
ATOM	918	N	HIS	139	. 14.787		-5.507	1.00	
MOTA	919	CA	HIS	139	15.305	9.227	-4.109	1.00	
ATOM	920	c	HIS	139	15.303	7.828	-3.508	1.00	
ATOM	921	ō	HIS	139	14.920	7.649	-2.347	1.00	
ATOM	922	CB	HIS	139	14.544	10.133	-3.130	1.00	
ATOM	923	CG	HIS	139	14.222	11.495	-3.664	1.00	
ATOM	924		HIS	139	15.148	12.292	-4.311	1.00	
ATOM	925		HIS	139	13.067	12.198	-3.645	1.00	
ATOM .	926		HIS	139	14.574	13.430	-4.666	1.00	
ATOM	927		HIS	139	13.308	13.397	-4.271	1.00	
ATOM	928	N.	SER	140	15.696	6.842	-4.305		
ATOM	929	CA	SER	140	15.753	5.455	-3.852	1.00 2	
ATOM	930	C	SER	140	17.193	5.111	-3.535	1.00 2	
MOTA	931	0	SER	140	18.123	5.586	-4.187	1.00 2	
ATOM	932	СВ	SER	140	15.225	4.514	-4.933	1.00 2	
MOTA	933	⊙G	SER	140	15.283	3.165	-4.519	1.00 3	
ATOM	934	N	GLU	141	17.374	4.288	-2.520	1.00 2	
ATOM	935	CA	GLU	141	18.704	3.893	-2.138	1.00 2	
ATOM	936	c	GLU	141	19.142	2.726		1.00 2	
ATOM	937	0	GLU	141	18.301	1.967		1.00 2	
MOTA	938.	CB	GLU	141	18.735	3.551		1.00 2	
MOTA	939	CG	GLU	141	17.921	4.524		1.00 3	

MOTA	94		D GL	U 141	18.5	78 4.85	5 1.55	1.00 69.28
MOTA	94		El GL		19.4			
MOTA	94		E2 GL		18.2	52 5.92		
MOTA	94							•
ATOM	94		A AL				4 -4.239	
ATOM	94				22.3	95 1.43	-3.643	
ATOM	94	_			23.1			
ATOM	94				21.1		7 -5.696	
ATOM	94				22.7		-3.521	
ATOM	94	_			24.0		-2.914	
ATOM	95				24.6		-3.709	1.00 24.43
ATOM	95				23.9			
ATOM	95				23.79			1.00 17.55
ATOM	95				22.8			1.00 18.42
atom Atom	954		D1 PHE		23.3			1.00 20.30
ATOM	955		D2 PHE		21.49			1.00 22.70
ATOM	956 957		E1 PHE		22.49			1.00 23.21
ATOM	958		E2 PHE		20.62			1.00 24.05
ATOM	959				21.13			1.00 22.11
ATOM	960		THR		25.92			1.00 20.15
ATOM	961		THR THR		26.59		-4.215	1.00 19.72
ATOM .	962		THR	144 144			-3.088	1.00 24.16
ATOM	963			144	27.57 27.79		-2.081	1.00 26.93
ATOM	964			144			-5.049	1.00 26.86
ATOM	965		2 THR	144	28.63 27.32		-4.242	1.00 28.49
ATOM	966		VAL	145	26.72		-6.228	1.00 29.29
ATOM	967			145	27.14		-3.178	1.00 17.10
ATOM	968		VAL	145	27.89	-	-2.134 -2.735	1.00 16.68
ATOM	969	O	VAL	145	28.039		-3.957	1.00 24.06 1.00 26.50
ATOM	970	СВ	VAL	145	25.944			1.00 20.66
ATOM	971	CG:	1 VAL	145	25.320		-0.516	1.00 20.88
ATOM	972	CG:	2 VAL	145	24.915		-2.175	1.00 19.97
ATOM	973	N	ASN	146	28.439		-1.882	1.00 19.01
ATOM	974	CA	ASN	146	29.158		-2.374	1.00 17.59
atom	975	С	ASN	146		-10.130	-2.342	1.00 20.14
ATOM	976	0	ASN	146		-10.694	-1.282	1.00 20.83
ATOM	977	CB	ASN	146	30.368	-9.305	-1.497	1.00 19.20
ATOM	978	CG	ASN	146		-10.590	-1.906	1.00 31.52
MOTA	979		ASN	146		-11.106	-3.005	1.00 32.16
ATOM	980		ASN	146		-11.134	-1.007	1.00 23.72
ATOM	981	N	PHE	147	27.647	-10.476	-3.503	1.00 16.06
ATOM	982	CA	PHE	147		-11.574	-3.564	1.00.16.26
ATOM	983	С	PHE	147		-12.918	-3.518	1.00 23.42
ntom Ntom	984	0	PHE	147		-13.954	-3.654	1.00 26.61
	985	CB	PHE	147		-11.471		1.00 17.81
ltom Ltom	986	CG.	PHE	147		-10.406		1.00 18.52
LTOM	987		PHE	147				1.00 20.66
TOM	988		PHE	147	24.891			1.00 19.12
TOM	989		PHE	147	22.621			1.00 20.73
TOM	990 991		PHE	147	23.909	-8.237		1.00 21.58
TOM	992	CZ	PHE	147	22.772		-4.532	1.00 19.82
TOM	993	N CA	GLY	148				.00 17.70
	223	-	GLY	148	29.453	-14.132	-3.241 1	.00 17.66

ATOM	99	4 C	GLY	148	29.147 -14.747	-1.88	4 1.00 23.36
ATOH		5 0	GLY	148	29.266 -15.959		2 1.00 23.54
MOTA		6 N	ASF	149	28.835 -13.887		
MOTA	99	7 CJ	A ASP	149	28.469 -14.333	0.42	
YTON	-99	8 C	ASP	149	26.958 -14.13 <u>4</u>	0.57	
MOTA	99		ASP	149	26.498 -13.190		
ATOH	100		ASP	149	29.220 -13.538	1.49	
ATOM	100				29.092 -14.150	2.880	1.00 20.27
MOTA	100		1 ASP		28.363 -15.132	3.086	
ATOM	100		2 ASP			3.791	1.00 25.46
MOTA	1004		THR		26.205 -15.062	0.001	
MOTA	1005				24.747 -15.040	0.013	1.00 18.86
MOTA	1006		THR		24.105 -14.621	1.338	1.00 21.45
ATOM	1007		THR	150	23.200 -13.784	1.363	1.00 20.49
ATOM	1008			150	24.195 -16.417	-0.405	1.00 24.26
ATOM	1009			150	24.591 -16.690	-1.752	1.00 32.55
MOTA	1010			150	22.684 -16.457	-0.293	1.00 19.05
MOTA	1011		GLU	151	24.601 -15.181	2.436	
ATOM	1012			151	24.082 -14.897	3.773	1.00 18.69
ATOM	1013		GLU	151	24.347 -13.469	4.195	1.00 22.02
ATOM ATOM	1014		GLU	151	23.470 -12.821	4.757	1.00 21.78
ATOM	1015	CB	GLU	151	24.692 -15.857	4.797	1.00 20.69
ATOM	1016	CG	GLU	151	23.943 -17.179	4.947	1.00 32.68
ATOM	1017 1018	CD	GLU	151	22.941 -17.153	6.089	1.00 52.10
ATOM	1019		GLU GLU	151	21.810 -16.675	5.911	1.00 56.21
ATOM	1020	N N		151	23.309 -17.631	7.187	1.00 41.59
ATOM	1020	CA	GLU GLU	152	25.568 -13.001	3.940	1.00 17.95
ATOM	1022	CA	GLU	152	25.977 -11.642	4.274	1.00 17.12
ATOM	1023	0	GLU	152 152	25.135 -10.625 24.758 -9.602	3.536	1.00 22.64
ATOM	1024	СВ		152	24.758 -9.602 27.461 -11.430	4.104	1.00 22.21
ATOM	1025		GLU	152	28.005 -10.055	3.958	1.00 17.92
MOTA	1026		GLU	152	27.855 -9.711	4.315	1.00 15.21
ATOM	1027		GLU	152	27.469 -10.580	5.778 6.595	1.00 19.13
ATOM	1028		GLU	152	28.127 -8.546	6.116	1.00 11.56
ATOM	1029	N	ALA	153	24.767 -10.957	2.306	1.00 14.61
ATOM	1030	CA	ALA	153	23.931 -10.079	1.508	1.00 19.83
ATOM	1031	c	ALA	153	22.571 -9.958	2.188	1.00 19.28 1.00 22.30
MOTA	1032	0	ALA	153	22.112 -8.843	2.431	1.00 22.30
MOTA	1033	CB	ALA	153	23:798 -10.601	0.066	1.00 19.78
MOTA	1034	N	LYS	154	21.980 -11.084	2.588	1.00 16.20
MOTA	1035	CA	LYS	154	20.677 -11:029	3.255	1.00 16.61
MOTA	1036	С	LYS	154	(20.759 -10.122	4.475	1.00 24.53
MOTA	1037	0	LYS	.154	19.916 -9.247	4.655	1.00 25.74
MOTA	1038	CB	LYS	154	20.212 -12.391	3.754	1.00 19.25
MOTA	1039	CG	LYS	154	19.848 -13.422	2.730	1.00 18.46
atom	1040	CD	LYS	154	19.395 -14.664	3.474	1.00 21.03
MOTA	1041	CE	LYS	154	19.798 -15.955	2.776	1.00 29.51
MOTA	1042	NZ	LYS	154	19.175 -17.133	3.456	1.00 41.79
MOTA	1043	N	LYS	155	21.741 -10.364		1.00 21.41
MOTA	1044		LYS	155	21.890 -9.542		1.00 21.15
ATOM	1045		LYS	155	22.018 -8.088		1.00 26.11
MOTA	1046		LYS	155	21.184 -7.253		1.00 27.67
NTOM	1047	CB	LYS	155	23.102 -9.974		1.00 22.36
							

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ATON	104B	∞	LYS	155	23.154	-9.290	8.732	1.00 45.02
HOTA	1049	CD	LYS	. 155	24.283	-9.774	9.598	1.00 62.54
ATON	1050	CE	LYS	155	24.327	-8.992	10.905	1.00 92.82
ATON	1051	NZ	LYS	155	25.397	-9.487	11.812	1.00110.00
ATON	1052	N	GLŅ	156	22.975		5.212	1.00 19.93
MOTA	1053	CX	GLŅ	.156	23.264	-6.521	4.679	1.00 19.33
MOTA	1054	С	GLN	156	22.015	-5.809	4.166	1.00 22.80
ATOH	1055	0	GLN	156	21.831	-4.601	4.363	1.00 23.72
ATON	1056	СВ	GLN	156	24.278	-6.652	3.543	1.00 21.20
ATOM	1057	CC	GLN	156	25.363	-5.589	3.538	1.00 57.09
ATOM	1058	CD	GLN	156	26.548	-5.987	2.678	1.00 93.87
ATOM	1059	OE	GLN	156	26.866	-7.171	2.529	1.00 94.48
ATOM	1060		GLN	156	27.216	-4.999	2.115	1.00 89.29
ATOM	1061	N	ILE	157	21.142	-6.585	3.543	1.00 17.92
ATOM	1062	CA	ILE	157	19.899	-6.082	2.970	1.00 18.11
MOTA	1063	c	ILE	. 157	18.868	-5.832	4.062	1.00 25.86
ATOM	1064	o	ILE		18.338	-4.723	4.203	1.00 27.68
ATOM	1065	СВ	ILE	157	19.330	-7.108	1.970	1.00 21.30
ATOM	1066		ILE	157	20.209	-7.151	0.716	1.00 21.24
ATOM	1067		ILE	157	17.871	-6.825	1.672	1.00 21.77
ATOM	1068		ILE	157	19.683	-8.024	-0.396	1.00 29.59
ATOM	1069	N	ASN	158	18.578		4.804	1.00 20.59
ATOM	1070	CA	ASN	158	17.632	-6.879	5.901	1.00 19.17
ATOM	1071	c	ASN	158	17.949	-5.804	6.941	1.00 23.91
ATOM	1072	ō	ASN	158	17.049	-5.178	7.498	1.00 25.85
ATOM	1073	CB	ASN	158	17.542	-8.284		1.00 15.15
ATOM	1074	CG	ASN	158	16.834	-9.240	5.549	1.00 28.18
ATOM	1075		ASN	158	15.934	-8.832	4.806	1.00 35.81
ATOM	1076		ASN	158		-10.506		1.00 16.31
ATOM	1077	N	ASP	159	19.230	-5.511	7.097	1.00 17.77
ATOM	1078	CA	ASP	159	19.682	-4.481	8.020	1.00 16.35
ATOM	1079	C	ASP	159	19.163	-3.153	7.484	1.00 23.88
ATOM	1080	0	ASP	159	18.614	-2.361	8.230	1.00 25.41
ATOM	1081	СВ	ASP	159	21.204	-4.418	8.066	1.00 17.14
ATOM	1082	CG	ASP	159	21.817	-5.525	8.903	1.00 24.66
ATOM	1083		ASP	159	21.088	-6.358	9.471	1.00 21.78
ATOM	1084		ASP	159	23.068	-5.554	8.994	1.00 34.61
ATOH	1085	N	TYR	160	19.309	-2.929	6.178	1.00 18.74
ATOM	1086	CA	TYR	160	. 18.845	-1.697	5.546	1.00 17.14
ATOM	1087	C	TYR	160	17.381	-1.434	5.876	1.00 21.00
ATOH	1088	0	TYR	160	17.038	-0.379	6.410	1.00 20.20
ATOM	1089	CB	TYR	160	19.051	-1.791	4.034	1.00 17.42
ATOM	1090	CG	TYR	160	18.250	-0.818	3.205	1.00 18.83
ATOM	1091		TYR	160	18.551	0.541	3.195	1.00 21.81
ATOM	1092		TYR	160	17.190	-1.261	2.416	1.00 19.83
ATOM	1093		TYR	160	17.815	1.432	2.420	1.00 24.46
MOTA	1094	CE2	TYR	160	16.449	-0.372	1.638	1.00 20.73
ATOH	1095	cz	TYR	160	16.770	0.970	1.642	1.00 30.99
MOTA	1096	ОН	TYR	160	16.064	1.847	0.853	1.00 40.24
ATOH	1097	N	VAL	161	16.537	-2.417	5.595	1.00 16.86
ATOM	1098	CA	VAL	161	15.113	-2.316	5.850	1.00 16.19
ATOM	1099	c c	VAL	161	14.841	-2.067	7.326	1.00 20.13
ATOM	1100	ō	VAL	161	14.189	-1.090	7.677	1.00 20.42
ATOM	1101	CB	VAL	161	14.392	-3.605	5.426	1.00 20.73

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ATOH	110)2 C	G1 VA	L 161	12.89	2 -3.49	2 5.67	7 1.00 19.90	
ATON)3 C	G2 VA	L 161					
ATOM	110	4 N	GL	J 162			_		
ATOM			A GL	J 162		2 -2.822			
ATOM		-	GL	J 162					
MOTA			GLU	J 162	. 14.71				
MOTA			-		16.01				
MOTA					15.80				
ATOM	111				16.58	5 -5.079			
ATOM	111		E1 GLU		16.75	6.127	11.809		
ATOM	111:		E2 GLU	•	17.009	-4.924	13.633		
ATOM	111:		LYS		16.757	-1.009	9.800		
ATOM	1114				17.286		10.145		
MOTA	1119		LYS		16.342	1.386		1.00 22.08	
ATOM	1110				16.012		10.332	1.00 22.56	
ATOM-					18.685		9.517	1.00 23.21	
ATOM	1118				19.222				
ATOM					20.483		8.401		
ATOM	1120			163	21.113		8.140		
ATOM	1121				22.310		7.249	1.00 90.37	
MOTA MOTA	1122		GLY	164	15.831	1.159	8.422		
ATOM	1123			164	14.939	2.107	7.785		
ATOM	112 4 1125		GLY	•	13.534		8.346	1.00 25.15	
ATOM	1125		GLY	164	12.854	3.127	8.272	1.00 27.19	
ATOM	1127		THR THR	165	13.102	0.985	8.910		
ATOM	1128		THR	165 165	11.766	0.861	9.492	1.00 18.76	
ATOM	1129	o	THR	165	11.944 11.008	0.902	11.005	1.00 26.62	
ATOM	1130			165	11.008	0.645	11.774	1.00 27.29	
ATOM	1131	OGI		165	11.893	-0.484 -1.582	9.116	1.00 12.69	
ATOM	1132	CG2		165	10.842	-0.585	9.557		
ATOM	1133	N	GLN	166	13.172	1.211 -	7.611 11.409	1.00 3.04	
ATOM	1134	CA	GLN	166	13.570	1.276	12.808	1.00 23.31	
MOTA	1135	С	GLN	166	13.075	0.102	13.664	1.00 22.33	
MOTA	1136	0	GLN	166	12,495	0.276	14.752	1.00 27.93 1.00 29.62	
MOTA	1137	CB	GLN	166	13.278	2.656	13.417	1.00 23.16	
ATOM	1138	CG	GLN	166	14.291	3.701	12.952	1.00 54.02	
MOTA	1139	CD	GLN	166	14.384	4.920	13.852	1.00 89.44	
MOTA	1140	OE1	GLN	166	13.958	4.907	15.011	1.00 87.60	
ATOM	1141	NE2	GLN	166	14.968	5.984	13.322	1.00 86.30	
MOTA	1142	N	GLY	167	13.350				
ATOM .	1143	CA	GLY	167	12.995	-2.305	13.872	1.00 17.73	
ATOM	1144	C	GLY	167	11.645		13.598	1.00 18.99	
MOTA	1145	0	GLY	167	11.356	-3.969	14.127	1.00 17.55	
ATOM	1146	N	LYS	168	10.814		12.791	1.00 18.38	
ATOM	1147	CA	LYS	168	9.497		12.507	1.00 20.25	
MOTA	1148	С	LYS	168	9.595		11.567	1.00 27.25	
MOTA	1149	0	LYS	168	8.819		11.667	1.00 30.06	
ATOM	1150	CB	LYS	168	8.529	-1.784		1.00 22.56	
MOTA	1151	CG	LYS	168		-2.206	12.103	1.00 39.22	
ATOM	1152	CD	LYS	168				1.00 58.14	
MOTA	1153	CE	LYS	168		-2.908	13.833	1.00 64.40	
MTOM.	1154	NZ	LYS	168			13.430	1.00 71.27	
ATOM	1155	N	ILE	169	10.517	-3.935	10.618	1.00 21.26	

MOTA	1156	CA	ILE	169	•	10.716	-5.031	9.682	1.00 19.37
MOTA	1157	C	ILE	169		12.096	-5.644	9.941	1.00 23.05
HOTA	1158	0	ILE	169		13.127	-4.966	9.921	1.00 24.91
MOTA	1159	CB	ILĖ	169		10.525	-4.573	8.210	1.00 21.73
ATOM	1160	CG1	ILE	169		9.060	-4.139	7.984	1.00 22.05
MOTA	1161	CG2	ILE	169		10.910	-5.699	7.258	1.00 20.93
ATOM	1162	CD1	ILE	169	. '	8.782	-3.495	6.634	1.00 17.20
ATOM	1163	N	VAL	170		12.096	-6.930	10,242	1.00 16.49
ATOM	1164	ÇA	VAL	170		13.321	-7.637	10.546	1.00 15.72
ATOM	1165	С	VAL	170		13.470	-8.824	9.597	1.00 19.76
ATOM	1166	0	VAL	170		12.478	-9.461	9.237	1.00 17.77
MOTA	1167	CB	VAL	170		13.279		11.998	1.00 20.38
ATOM	1168	CG1	VAL	170		13.353	-6.848	12.905	1.00 19.80
ATOM	1169	CG2	VAL	170		~11.979	-8.823	12.259	1.00 21.00
ATOM	1170	N	ASP	171		14.694	-9.035	9.114	1.00 19.34
ATOM	1171	CA	ASP	171			-10.112	8.162	1.00 20.38
ATOM	1172	С	ASP	171			-10.430	7.144	1.00 22.67
MOTA	1173	0	ASP	171			-11.473	7.179	1.00 23.02
ATOM	1174	CB	ASP	171			-11.359	8.832	1.00 23.70
ATOM	1175	CG	ASP	171		14.674	-12.312	9.480	1.00 42.06
MOTA	1176		ASP	171		13.524	-11.929	9.772	1.00 48.32
ATOM	1177		ASP	171		15.067	-13.473	9.715	1.00 45.87
ATOM	1178	N	LEU	172		13.731	-9.474	6.241	1.00 18.15
ATOM	1179		LEU	172	*	12.723	-9.547	5.196	1.00 17.36
ATOM	1180	C	LEU	172		12.976	-10.702	4.250	1.00 24.59
ATOM	1181	ō	LEU	172		12.062	-11.465	3.915	1.00 26.72
ATOM	1182	СВ	LEU .	172		12.715	-8.238	4.413	1.00 16.47
MOTA	1183	CG	LEU	172	٠.	11.669	-8.014	3.338	1.00 20.87
ATOM	1184		LEU	172	٠.	10.307	-8.399	3.853	1.00 20.88
ATOM	1185		LEU	172		11.704	-6.568	2.908	1.00 21.64
MOTA	1186	N	VAL	173		14.227	-10.804	3.812	1.00 19.02
MOTA	1187	CA	VAL	173		14.662	-11.837	2.892	1.00 16.81
ATOM	1188	Ç.	VAL	173			-13.031	3.701	1.00 20.99
MOTA	1189	0	VAL	173		16.010	-12.901	4.555	1.00 22.14
ATOM	1190	CB	VAL	173			-11.312	1.970	1.00 19.26
ATOM	1191	CG1	VAL	173			-12.402	1.023	1.00 19.04
ATOM	1192		VAL	173			-10.109	1.190	1.00 18.18
ATOM .	1193	N	LYS	174			-14.179	3.441	1.00 17.06
ATOM	1194	CA	LYS	174			-15.430	4.118	1.00 17.44
ATOM	1195	С	LYS	174			-16.268	3.331	1.00 25.16
ATOM	1196	0	LYS	174			-17.274	3.841	1.00 27.74
ATOM	1197	CB	LYS	174			-16.227	4.316	1.00 20.15
ATOM	1198	CG	LYS	174		12.400	-15.414	4.914	1.00 35.14
ATOM	1199	CD	LYS	174		12.437	-15.391	6.423	1.00 47.19
ATOM	1200	CE	LYS	174		11.459	-14.373	6.961	1.00 58.72
MOTA	1201	NZ	LYS	174			-14.807	8.225	1.00 76.00
MOTA	1202	N	GLU	175			-15.900	2.074	1.00 19.67
MOTA	1203	CA	GLU	175			-16.614	1.229	1.00 19.84
ATOM	1204	С	GLU	175			-16.011	-0.162	1.00 26.26
MOTA	1205	0	GLU	175			-15.359	-0.678	1.00 27.25
MOTA	1206	CB	GLU	175			-18.111	1.131	1.00 21.25
MOTA	1207	CG	GĻU	175			-18.481	0.674	1.00 38.11
MOTA	1208	CD	GLU	175			-19.975	0.813	1.00 78.80
MOTA	1209	OE1	GLU	175		15.785	-20.646	1.555	1.00 83.96

ATOM	. 121	0 0	E2 GLU	175	14.066 -20.476 0.199 1.00 82.02
MOTA	121	1 - N	LEU	176	18.398 -16.215 -0.749 1.00 20.61
MOTA	121	2 C			18.719 -15.688 -2.076 1.00 18.97
MOTA	121	3 C	LEU	176	19.145 -16.786 -3.038 1.00 23.08
ATOM	. 121	4 0	LEU	176	19.766 -17.779 -2.637 1.00 21.51
_ ATOM	. 121	5 C1	B LEU	176	19.851 -14.657 -2:005 1.00 19.21
MOTA	1216	S CC	LEU	176	19.592 -13.270 -1.414 1.00 25.41
ATOM	1217	Ĉ CI	Ol LEU	1,76	20.909 -12.527 -1.314 1.00 24.90
MOTA	1218	CI	2 LEU	176	18.613 -12.504 -2.287 1.00 31.01
MOTA	1219	N	ASP	177	18.817 -16.595 -4.315 1.00 21.83
ATOM	1220		ASP	177	19.174 -17.555 -5.370 1.00 21.94
ATOM	1221	. c	ASP	177	20.675 -17.647 -5.524 1.00 28.47
ATOM	1222		ASP	177	21.414 -16.775 -5.070 1.00 31.66
ATOM	1223			177	18.545 -17.182 -6.714 1.00 24.78
ATOM	1224			177	17.025 -17.142 -6.655 1.00 50.17
ATOM	1225		1 ASP		16.363 -18.193 -6.746 1.00 51.41
ATOM	1226		2 ASP	177	16.483 -16.020 -6.513 1.00 63.80
ATOM	- 1227		ARG	178	21.103 -18.679 -6.236 1.00 22.85
MOTA	1228			178	22.522 -18.958 -6.470 1.00 21.85
MOTA	1229		ARG	178	23.191 -17.927 -7.386 1.00 24.51
ATOM	1230	-	ARG	178	24.348 -17.569 -7.189 1.00 26.94
ATOM	1231			178 .	
ATOM	1232		ARG		21.805 -21.462 -6.360 1.00 36.59
ATOM	1233	CD	ARG	178	20.349 -21.385 -6.826 1.00 61.75
MOTA MOTA	1234	NE	ARG	178	19.426 -22.048 -5.902 1.00 86.02
ATOM	1235	cz	ARG	178	18.120 -21.795 -5.819 1.00109.62
ATOM	1236 1237		LARG	178	17.556 -20.877 -6.589 1.00106.33
ATOM	1237	NA.	ARG	178	17.364 -22.485 -4.972 1.00 95.24
ATOM	1239	CA	ASP ASP	179 179	22.444 -17.422 -8.355 1.00 19.98
ATOM	1240	CV	ASP	179	22.946 -16.433 -9.310 1.00 20.64
ATOM	1241	ō	ASP	179	22.460 -15.014 -8.999 1.00 26.46
ATOM	1242	СВ	ASP	179	22.314 -14.174 -9.896 1.00 27.73 22.532 -16.813 -10.737 1.00 24.57
ATOM	1243	CG	ASP	179	22.532 -16.813 -10.737 1.00 24.57 21.033 -17.092 -10.856 1.00 59.63
ATOM	1244		ASP	179	20.259 -16.796 -9.918 1.00 65.53
ATOM	1245		ASP	179	20.639 -17.624 -11.919 1.00 71.83
ATOM	1246	N	THR	180	22.233 -14.739 -7.729 1.00 71.83
ATOM	1247	CA	THR	180	21.760 -13.426 -7.364 1.00 20.37
ATOM	1248	c	THR	180	22.871 -12.388 -7.545 1.00 23.03
ATOM	1249	0	THR	180	23.967 -12.536 -6.998 1.00 22.71
ATOM	1250	СВ	THR	180	21.225 -13.396 -5.915 1.00 32.99
ATOM	1251	OG1	THR	180	20.180 -14.370 -5.763 1.00 31.89
MOTA	1252		THR	180	20.663 -12.024 -5.589 1.00 34.35
MOTA	1253	N	VAL	181	22.607 -11.385 -8.383 1.00 18.85
ATOM	1254	CA	VAL	181	23.554 -10.297 -8.634 1.00 18.00
MOTA	1255	C 1	VAL	181	22.866 -8.934 -8.541 1.00 21.23
MOTA	1256	0	VAL	181	23.540 -7.901 -8.499 1.00 22.38
MOTA	1257	CB	VAL	181	24.245 -10.404 -10.022 1.00 21.67
MOTA	1258		VAL	181	25.156 -11.615 -10.072 1.00 21.73
MOTA	1259	CG2	VAL	181	23.209 -10.447 -11.136 1.00 21.58
MOTA	1260	N	PHE	182	21.537 -8.943 -8.454 1.00 15.29
ATOM	1261	CA	PHE	182	20.732 -7.723 -8.378 1.00 14.49
ATOM	1262	С	PHE	182	19.496 -8.014 -7.527 1.00 20.22
ATOM	1263	0	PHE	182	18.733 -8.929 -7.838 1.00 20.58
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ATO		4 CB	PHE 182	. 20.3	37 -7.3	15 .0 00	
ATO	M 126		PHE 182	19.5			
ATO	M 126			19.5			
ATO	M 126			18.9			
ATO		8 CE1 1		18.8		B2 -11.09	
ATO							
ATO			HE 182	18.2		35 -11.21	
ATO			LA 183	18.10		8 -10.15	
ATOR			LA 183	19.3			
ATON			LA 183	18.20			
ATOH			LA 183	17.46			
ATOH			LA 183	18.07			
ATOM				18.65			
MOTA	•	_		16.13	=		1.00 16.09
ATOM				15.26			
ATOM				14.43			
ATOH				13.74			1.00 17.17
ATOM				14.33			1.00 16.99
ATOM				13.43			1.00 22.65
ATOM	1283			12.97			1.00 21.80
MOTA	1284	CD2 LE		12.22			1.00 28.05
ATOM	1285	N VA	-	14.492			1.00 14.81
ATOM		CA VA		13.72			1.00 16.22
ATOM	1286	C VA		12.849		-1.363	1.00 18.85
MOTA	1287	O VA		13.344			1.00 16.95
ATOM	1288	CB VA		14.641			1.00 20.90
ATOM		CG1 VA				0.971	1.00 21.00
ATOH	1290	CG2 VA		15.433		-0.496	1.00 20.06
ATOM	1291	N ASI		11.541		-1.382	1.00 16.14
ATOM	1292	CA ASI		10.551	-2.427	-1.292	1.00 16.68
ATOM	1293	C ASN		9.653		-0.083	1.00 21.69
ATOM	1294	O ASA		9.051	-3.733	0.052	1.00 23.15
ATOM	1295	CB ASN		9.734	-2.398	-2.595	1.00 12.94
ATOM	1296	CG ASN		8.838	-1.204	-2.696	1.00 19.95
ATOM	1297	OD1 ASN		7.664	-1.250	-2.325	1.00 11.91
ATOM	1298	ND2 ASN		9.391	-0.106		1.00 10.24
ATOM	1299	N TYR		9.578	-1.687		1.00 15.75
ATOM	1300	CA TYR		8.773	-1.806		1.00 14.62
ATOM		C TYR		8.078	-0.471	2.290	1.00 22.42
ATOM		O TYR	187	8.484	0.581	1.772	1.00 23.11
ATOM		CB TYR	187	9.676	-2.184		1.00 15.24
		CG TYR	187	10.667	-1.084		1.00 18.84
ATOM ATOM		CD1 TYR	187	10.268	0.007	4.335	.00 21.93
	1306	CD2 TYR	187	12.004	-1.132		.00 19.78
ATOM ATOM		CEI TYR	187	11.171	1.013		.00 23.73
		CE2 TYR	187	12.911	-0.128		.00 20.67
ATOM		CZ TYR	187	12.484	0.935		.00 28.94
ATOM		OH TYR	187	13.374	1.906		.00 29.27
ATOM		N ILE	188	7.058	-0.517		.00 19.02
ATOM		CA ILE	188	6.304	0.673		.00 18.03
ATOM		ILE	188	5.626	0.411		.00 23.51
ATOM		. ILE	188	4.922	-0.581		.00 25.66
ATOM		B ILE	188	5.238	1.063		.00 19.71
ATOM		G1 ILE	188	4.464	2.302		.00 19.71
MOTA	1317 C	G2 ILE	188		-0.091		.00 16.85
		•				J J.	10.65

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ATOM	1318	CD1	ILE	188	3.574	2.900	1.869	1.00 28.04
ATOM	1319	N	PHE	189	5. 9 10	1.256	. 5.833	1.00 18.63
MOTA	1320	CA	PHE	189	5.309	1.119	•	1.00 17.30
MOTA	1321	С	PHE	189	4.489	2.368	7,452	1.00 18.72
MOTA	1322	0	PHE	189	5.002	3.490	7.368	1.00 16.24
ATOM	1323	CB	PHE	189	6.383	0.922	8.217	1.00 19.37
MOTA	1324	CG	PHE	189	5.850	1.018	9.616	1.00 21.45
ATOM	1325	CD1	PHE	. 189	5.200	-0.063	10.199	1.00 23.96
ATOM	1326	CD2	PHE	189	5.930	2.213	10.327	1.00 23.47
MOTA	1327	CEL	PHE	189	4.637	0.048	11.459	1.00 24.52
MOTA	1328	CE2	PHE	189	5.366	2.319	11.586	1.00 25.92
MOTA	1329	CZ	PHE	189	4.719	1.237	12.149	1.00 23.37
MOTA	1330	N ·	PHE	190	3.222	2.168	7807	1.00 15.59
MOTA	1331	CA	PHE	190	2.340	3.275	8.123	1.00 15.41
ATOM	1332	С	PHE	190	1.589	3.073	9.425	1.00 21.53
ATOM	1333	0	PHE	190	1.228	1.957	9.784	1.00 22.76
ATOM	1334	CB	PHE	190	1.335	3.500	7.002	1.00 17.31
ATOM	1335	CG	PHE	190	0.448	4.690	7.222	1.00 18:57
ATOM	1336	CD1	PHE	190	0.996	5.967	7.286	1.00 21.44
ATOM	1337	CD2	-	190	-0.927	4.543	7.384	1.00 19.70
ATOM	1338	CE1		190	0.193	7.067	7.504	1.00 22.32
ATOM	1339	CE2	PHE		-1.733	5.647	7.602	1.00 22.76
ATOM	1340	cz	PHE	190	-1.177	6.904	7.662	1.00 21.21
ATOM	1341	N	LYS	191	1.369	4.171	10.130	1.00 19.61
ATOM	1342	CA	LYS	191	0.640	4.168	11.386	1.00 20.03
ATOM	1343	c	LYS		0.261	5.617	11.631	1.00 23.60
ATOM	1344	ō	LYS	191	0.952	6.344	12.344	1.00 24.22
ATOM	1345	СВ	LYS	191	1.514	3.650	12.525	1.00 24.39
ATOM	1346	CG	LYS	191	0.776	3.575	13.844	1.00 51.75
ATOM	1347	CD	LYS	191	1.726	3.537	15.032	1.00 59.10
ATOM	1348	CE	LYS	191	0.934	3.604	16.307	1.00 60.54
ATOM	1349	NZ		191	0.090	4.820	16.277	1.00 61.59
ATOM	1350	N	GLY	192	-0.751	6.065	10.899	1.00 18.83
ATOM	1351	CA	GLY	192	-1.207	7.431	11.028	1.00 17.89
MOTA	1352	С	GLY	192	-2.017	7.706	12.270	1.00 21.28
ATOM	1353	0	GLY	192	-2.533	6.800	12.925	1.00 21.21
MOTA	1354	N	LYS	193	-2.085	8.980	12.620	1.00 18.93
ATOM	1355	CA	LYS	193	-2.840	9.414	13.779	1.00 20.11
MOTA	1356	C	LYS	193	-4.170	9.929	13.239	1.00 28.57
MOTA	1357	Ο.	LYS	193	-4.200	10.621	12.222	1.00 30.68
MOTA	1358	СВ	LYS	193	-2.066	10.518	14.492	1.00 22.77
ATOM	1359	CG	LYS	193	-0.605	10.152	14.635	1.00 59.22
ATOM	1360	CD	LYS	193	0.255	11.289	15.098	1.00 85.61
ATOM	1361	CE	LYS	193	1.716	10.864	15.104	1.00104.77
ATOM	1362	NZ	LYS	193	2.595	11.990	15.496	1.00110.00
MOTA	1363	N	TRP	194	-5.275	9.502	13.842	1.00 22.28
ATOM	1364	CA	TRP	194	-6.580	9.956	13.389	1.00 19.23
ATOM	1365	C	TRP	194	-6.729	11.449	13.600	1.00 22.08
MOTA	1366	0	TRP	194	-6.345	11.975	14.642	1.00 20.61
ATOM	1367	CB	TRP	194	-7.709	9.232	14.127	1.00 16.80
ATOM	1368	CG	TRP	194	-7.820	7.791	13.793	1.00 17.43
ATOM	1369		TRP	194	-7.806	6.739	14.660	1.00 20.34
MOTA	1370		TRP	194	-7.894	7.235	12.478	1.00 17.41
ATOM	1371		TRP	194	-7.850	5.558	13.962	1.00 20.08

ATOM	1372	•	TRP	194	-7.920		12.633	1.00 21.75
MOTA	1373	CE:		194	-7.961		-	
MOTA	1374	CZZ		194	-7.969		11.530	1.00 20.89
MOTA	1375	CZ3	TRP	194	-8.009	6.933	10.111	1.00 20.61
ATOM	1376	CH2	TRP	194	-8.028		10.286	1.00 21.45
ATOM	1377	N	GLU	195	-7.308	12.105	12.578	1.00 18.47
MOTA	1378	CA	GĽU	195	-7.643	13.509	12.721	1.00 17.31
ATOM	1379	C:	GLU	195	-8.716	13.627	13.767	1.00 24.91
ATOM	1380	0	GLU	195	-8.750	14.601	14.501	1.00 26.50
MOTA	1381	CB	GLU	195	-8.193	13.966	11.359	1.00 17.55
MOTA	1382	CG	GLU	195	-8.065	15.493	11.231	1.00 17.56
MOTA	1383	CD	GLU	195	-8.452	15.898	9.839	1.00 24.67
ATOM	1384	OE1	GLU	195	-9.680	15.929	9.553	1.00 8.00
MOTA	1385	-	GLU	195	-7.532	16.193	9.032	1.00 29.08
ATOM	1386	N	ARG	- 196	-9.582	12.597	13.838	1.00 20.95
ATOM	1387	CA	ARG	196	-10.576	12.563	14.894	1.00 19.29
ATOM	1388	C	ARG	196	-10.401	11.246	15.606	1.00 19.82
ATOM	1389	ō	ARG	196	-10.955	10.248	15.172	1.00 17.50
MOTA	1390	СВ	ARG	196	-11.967	12.693	14.247	1.00 17.74
ATOM	1391	CG	ARG	196	-12.101	14.095	13.626	1.00 28.49
ATOM	1392	CD	ARG	196	-13.389	14.170	12.788	1.00 44.06
ATOM	1393	NE	ARG	196	-13.398	15.427	12.063	1.00 68.61
ATOM	1394	CZ	ARG	196	-14.509	15.946	11.626	1.00 90.69
ATOM	1395	NH1		196	-15.650	15.352	11.819	1.00 78.16
ATOM	1396			196	-14.475	17.076	10.984	1.00 79.20
ATOM	1397	N	PRO	197	-9.608	11.228	16.701	1.00 16.16
ATOM	1398	CA	PRO	197	-9.262	9.981	17.351	1.00 14.33
ATOM	1399	C	PRO	197	-10.380	9.296	18.090	
ATOM	1400		PRO	197	-11.441	9.847	18.328	1.00 19.66
ATOM	1401	СВ	PRO	197	-8.165	10.412	18.347	1.00 19.00
ATOM	1401		PRO	197	-8.160	11.956	18.384	1.00 22.80
ATOM	1403				-9.018	12.450	17.203	1.00 22.80
ATOM	1404	CD N	PRO	197	-10.067	8.035	18.439	1.00 15.54
ATOM	1405	CA	PHE PHE	198 198	-10.067	7.199	19.238	1.00 15.34
ATOM	1406	C			-10.338	7.325	20.641	1.00 25.44
ATOM	1400	0	PHE PHE	198 198	-9.150	7.325	20.823	1.00 25.44
ATOM	1407	CB				7.366 5.717		
ATOM			PHE	198	-10.867		18.834	1.00 16.18
ATOM	1409 1410	CG	PHE	198	-11.560	5.392	17.553	1.00 18.18
		CD1	PHE	198	-12.931	5.214	17.519	1.00 22.32
ATOM	1411	CD2 CE1	PHE	198	-10.847	5.303 4.958	16.366	
	1412	_		198 198	-13.589 -11.490	5.049	16.321 15.172	1.00 23.71 1.00 23.38
ATOM	1413	CE2						
ATOM	1414	cz	PHE	198	-12.866	4.877	15.146	1.00 21.79
MOTA	1415		GLU	199	-11.268	7.362	21.648	1.00 21.15
	1416	CA	GLU	199	-10.759	7.396	23.004	1.00 21.66
MOTA	1417	С	GLU	199	-10.458	6.001	23.469	1.00 25.73
ATOM -	1418	0	GLU	199	-11.331	5.146	23.467	1.00 27.78
ATOM	1419	CB	GLU	199	-11.738	8.127	23.941	1.00 24.14
MOTA	1420	CG	GLU	199	-11.845	9.597	23.490	1.00 44.18
ATOM	1421		GLU	199	-12.328	10.519	24.577	1.00 83.32
MOTA	1422	OE1		199	-12.652	10.035	25.695	1.00 77.92
MOTA	1423	OE2		199	-12.381		24.309	1.00 89.79
MOTA	1424	N		200	-9.187		23.862	1.00 18.99
MOTA	1425	CA	VAL	200	-8.765	4.484	24.356	1.00 17.54

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HOTA	1426	C	VAL	200	-9.690	3.888	25.421		21.63
HOTA	1427	0	VAL	200	-9.879	2.676	25.477		22.09
MOTA	1428	CB	VAL	200	-7.319	4.539	24.927	1.00	21.35
MOTA	1429	CG1	VAL	200	-6.796	3.147	25.174		21.50
MOTA	1430	CG2	VAL	200	-6.395	5.278	23.981	1.00	21.61
MOTA	.1431	N	LYS	201	-10.319	4.742	26.217		19.04
MOTA	1432	CA	LYS	201	-11.192	4.262	27.275	1.00	18.50
MOTA	1433	Ċ	LYS		-12.267	3.344	26.726	1.00	26.40
ATOM	1434	0	LYS		-12.581	2.335	27.342	1.00	26.71
ATOM	1435	СВ	LYS		-11.833	5.430	28.023	1.00	18.67
ATOH	1436	CG	LYS	201	-12.888	6.216	27.249	1.00	18.75
HOTA	1437	CD	LYS	201	-13.518	7.283	28.119	1.00	18.80
ATOM	1438	CE	LYS	201	-14.672	7.970	27.427	1.00	32.20
MOTA	1439	NZ	LYS	201	-15.326	8.972	28.307	1.00	45.27
ATOM	1440	N	ASP	202	-12.780	3.682	25.544	1.00	25.30
ATOM	1441	CA	ASP	202.	-13.834	2.923	24.899	<1.00 ·	26.03
MOTA	1442	Ç	ASP	202	-13.386	1.698	24:117	1.00	30.85
ATOM	1443	0	ASP	202	-14.216	0.984	23.546	1.00	30.98
ATOM	1444	СВ	ASP	202	-14.654	3.841	23.990	1.00	28.26
ATOM	1445	CG	ASP	202	-15.464	4.848	24.770	1.00	43.04
MOTA	1446	OD1	ASP	202	-16.258	4.401	25.628	1.00	52.14
ATOM	1447	OD2	ASP	202	-15.303	6.060	24.547	1.00	44.22
ATOM	1448	N	THR	203	-12.086	1.451	24.069	1.00	25.69
ATOM	1449	CA	THR	203	-11.587	0.297	23.347	1.00	24.08
ATOM	1450	C	THR	203	-11.707	-0.967	24.179	1.00	28.24
ATOM	1451	0	THR	203	-10.951	-1.189	25.130	1.00	29.39
MOTA	1452	CB	THR	203	-10.145	0.504	22.869	1.00	22.59
ATOM	1453	0G1	THR	203	-10.098	1.685	22.067	1.00	28.88
MOTA	1454	CG2	THR	203	-9.678	-0.665	22.029		18.61
MOTA	1455	N	GLU	204	-12.760	-1.722	23.891		23.91
MOTA	1456	. CA	GLU	204	-13.028	-2.986	24.560		24.34
ATOM	1457	С	GLU	204	-12.810	-4.070	23.516		27.16
ATOM	1458	0	GLU	204	-12.856	-3.820	22.309		26.99
ATOM	1459	CB	GLU	204	-14.483	-3.091	25.017		26.97
MOTA	1460	CG	GLU	204	-15.013	-2.054	25.993	1.00	
ATOM	1461	CD	GLU	204	-16.540	-2.055	26.029	1.00	
MOTA	1462	0E1	GLU	204	-17.155	-3.104	25.739	1.00	
MOTA	1463	OE2	GLU	204	-17.127	-0.993	26.318	1.00	
ATOM	1464	N	GLU		-12.598	-5.287	23.987	1.00	
MOTA	1465	CA	GLU	205	-12.393	-6.423	23.111	1.00	
MOTA	1466	С	GLU	205	-13.775	-6.929	22.733	1.00	
MOTA	1467	0	GLU	205	-14.668	-6.957	23.574	1.00	
ATOM	1468	CB	GLU	205	-11.610	-7.503	23.850	1.00	
MOTA	1469	CG	GLU	205	-11.100	-8.624	22.991	1.00	
MOTA	1470	CD	GLU	205	-10.167	-9.514	23.763	1.00	
MOTA	1471	OE1		205		-10.332	24.558	1.00	
ATOM	1472	OE2		205	-8.939	-9.387	23.610	1.00	
ATOH	1473	N	GLU	206	-13.966	-7.296	21.471	1.00 2	
ATOM	1474	CA	GLU	206	-15.267	-7.780 -8.773	21.021	1.00 2	
ATOM	1475	С	GLU	206	-15.158 -14.068	-8.773 -9.020	19.859 19.340	1.00 2	
ATOM	1476	0	GLU	206	-14.088	-6.603	20.673	1.00 2	
ATOH	1477	CB	GLU	206	-16.196	-5.848	21.889	1.00 4	
MOTA	1478	CG	GLU	206	-16.778	-4.339	21.667	1.00 7	
MOTA	1479	CD	GLU	206	-10.308	337	41.00/	1.00	5.50

ATOM	1480	OE	1 GLU	206	-17.346 -3.9	15 20.577	1.00 98.65
ATOM	. 1481	OE	2 GLU	206	-16.590 -3.5	76 22.614	1.00 53.79
ATOH	1482	N	ASP	207	-16.302 -9.3	56. 19.496	1.00 21.57
ATOM	1483	CA	ASP	207	-16.437 -10.3		
MOTA	1484	C	ASP	207	-16.049 -9.8	96 17.030	1.00 27.25
ATOM	1485	0	ASP	207	-16.278 -8.7	48 16.652	1.00 29.30
ATOM	1486	CB	ASP	207	-17.877 -10.8	85 18.367	1.00 22.21
ATOM	1487	CC	ASP	207	-18.238 -11.7		1.00 21.70
MOTA	. 1488		1 ASP	207	-17.788 -11.5		1.00 16.73
MOTA	1489	OD	2 ASP	207	-19.024 -12.7		1.00 33.68
ATOM	1490	N	PHE		-15.475 -10.8		1.00 21.30
ATOM	1491	CA	PHE	208	-15.066 -10.5		1.00 19.76
MOTA	1492	C	PHE	208	-15.395 -11.83		1:00 22.95
MOTA	1493	0	PHE	208	-14.783 -12.80		1.00 22.62
ATOM	1494	CB	PHE	208	-13.577 -10.23		1.00 21.53
ATOM	1495	CG	PHE	208	-13.176 -9.73		1.00 23.92
MOTA	1496	CD:		208	-13.071 -10.58		1.00 28.21
MOTA	1497		2 PHE	208	-12.948 -8.36		1.00 27.58
ATOM	1498		L PHE	208	-12.747 -10.11		
MOTA	1499		PHE	208	-12.624 -7.88		1.00 28.80
ATOM	1500	CZ	PHE	208	-12.525 -8.76		1.00 29.25
MOTA	1501	N		209	-16.413 -11.68		1.00 20.52
MOTA	1502	CA	HIS	209	-16.875 -12.76		1.00 20.85
ATOM	1503	С	HIS	209	-15.919 -13.20		1.00 23.39
MOTA	1504	.0	HIS	209	-15.855 -12.57		1.00 24.89
MOTA	1505	СВ	HIS	209	-18.212 -12.39		1.00 22.94
MOTA	1506	CG	HIS	209	-19.393 -12.86		1.00 27.59
MOTA	1507	ND1		209 .	-20.195 -13.90		1.00 30.09
ATOM	1508	-	HIS	209	-19.871 -12.47		1.00 29.90
MOTA	1509		HIS	209	-21.115 -14.14		1.00 29.82
MOTA MOTA	1510		HIS	209	-20.940 -13.29 -15.185 -14.28		1.00 30.00
ATOM	1511 1512	N CA	VAL	210 210	-14.271 -14.83		1.00 17.01
ATOM	1513	C.	VAL	210	-15.076 -15.76		1.00 22.79
ATOM	1514	0	VAL	210	-14.609 -16.15		1.00 24.05
ATOM	1515	СВ	VAL	210	-13.113 -15.60		1.00 15.44
ATOM	1516		VAL	210	-12.136 -14.61		1.00 14.45
ATOM	1517	CG2		210	-13.639 -16.55		1.00 15.19
ATOM	1518	N	ASP	211	-16.282 -16.12		1.00 21.95
MOTA	1519	CA	ASP	211	-17.230 -16.99		1.00 23.32
ATOM	1520	C	ASP	211	-18.628 -16.42		1.00 30.30
ATOM	1521	0	ASP	211	-18.818 -15.50		1.00 30.53
MOTA	1522	СВ	ASP	211	-17.226 -18.41		1.00 25.50
MOTA	1523	CG	ASP .	211	-16.125 -19.266		1.00 44.95
MOTA	1524		ASP	211	-14.933 -18.996		1.00 48.54
MOTA	1525	OD2	ASP	211	-16.463 -20.274		1.00 52.49
MOTA	1526	N	GLN	212	-19.622 -17.017		1.00 29.03
MOTA	1527	CA	GLN	212	-21.007 -16.581		1.00 29.12
ATOM	1528	С	GLN	212	-21.484 -17.190		1.00 34.30
MOTA	1529	0	GLN	212	-22.502 -16.770		1.00 36.41
MOTA	1530	CB	GLN	212	-21.918 -17.103	8.026	1.00 30.36
MOTA	1531	CĠ	GLN	212	-21.633 -16.563	6.630	1.00 49.34
MOTA	1532	CD .	GLN	212	-22.656 -15.539	6.172	1.00 64.74
MOTA	1533	0É1	GLN	212	-23.716 -15.377	6.779	1.00 59.48

MOTA	1534		2 GLN	212		-14.839		1.00 56.91
ATOH	1535	N	VAL	213		-18.166		
MOTA	1536	CA	VAL	213	- - · ·	-18.876		1.00 25.98
ATOM	1537	С	VAL	213		-18.912		1.00 26.83
HOTA	1538	.0	VAL	213		-19.375		1.00 24.65
ATOH	1539	CB	VAL	213	•	-20.338	-	1.00 28.98
MOTA	1540		l VAL			-20.351		1.00 28.50
ATOH	1541	CG	2 VAL	213	-20.327	-21.109	11.243	1.00 28.77
MOTA	1542	N	THR	214	-18.784	-18.409	12.851	1.00 23.26
HOTA	1543	CA	THR	214		-18.411	13.747	1.00 23.15
HOTA	1544	C	THR	214	-17.196	-16.989	14.050	1.00 27.67
MOTA	1545	0	THR	214	-17.353	-16.090	13.221	1.00 27.31
MOTA	1546	CB	THR	214	-16.461	-19.186	13.136	1.00 31.51
MOTA	1547	OG	THR	214	-16.968	-20.276	12.347	1.00 28.47
MOTA	1548	CG	THR	214	-15.551	-19.727	14.227	1.00 32.93
MOTA	. 1549	N	THR	215	-16.665	-16.793	15.250	1.00 25.25
MOTA	1550	CA	THR	215	-16.198	-15.496	15.694	1.00 25.06
ATOM -	1551	C	THR	215	-14.836	-15.607	16.353	1.00 26.60
ATOM	1552	0	THR	215		-16.697	16.655	1.00 26.58
ATOM	1553	CB	THR	215	-17.187	-14.856	16.708	1.00 39.80
ATOM	1554	0G1		215	-17.476	-15.787	17.760	1.00 40.04
MOTA	1555	CG2	THR		18.480	-14.468	16.029	1.00 43.19
ATOM	1556	N	VAL	216		-14.458	16.537	1.00 21.12
ATOM	1557	CA	VAL	216	-12.907	-14.343	17.166	1.00 19.03
ATOM	1558	C	VAL	216	-12.933		17.890	1.00 26.31
ATOM	1559	Ō	VAL	216	-13.723		17.550	1.00 26.92
ATOM	1560	СВ	VAL	216	-11.782		16.122	1.00 20.18
ATOM	1561		VAL	216	-10.557		16.590	1.00 19.92
ATOM	1562	CG2		216	-11.434		15.873	1.00 19.39
ATOM	1563	N:	LYS	217	-12.159		18.962	1.00 23.35
ATOM	1564	CA	LYS	217	-12.119		19.734	1.00 23.09
ATOM	1565	c	LYS	217.	-10.944		19.295	1.00 24.17
ATOM	1566	0	LYS	217		-11.320	19.004	1.00 21.99
ATOM	1567	CB	LYS	217	-12.028		21.226	1.00 27.15
ATOM	1568	CG	LYS	217	-13.130		21.712	1.00 34.17
ATOM	1569	CD	LYS	217	-14.487		21.553	1.00 37.75
ATOM		CE	LYS	217	-15.602		21.705	1.00 51.12
ATOM	1571	NZ	LYS	217	-15.661		20.501	1.00 59.22
ATOM	1572	N	VAL	218	-11.194	-9.509	19.216	1.00 20.37
ATOM	1573	CA	VAL	218	-10.183	-8.538	18.823	1.00 19.85
ATOM	1574	c	VAL	218	-10.486	-7.236	19.547	1.00 19.60
ATOM	1575	ō	VAL	218	-11.622	-7.030	19.985	1.00 19.43
ATOM	1576		VAL	218	-10.225	-8.280	17.295	1.00 25.31
ATOM	1577		VAL	218	-9.764	-9.521	16.540	1.00 26.00
ATOM	1578		VAL	218	-11.625	-7.885	16.853	1.00 24.90
ATOH	1579	N	PRO	219	-9.457	-6.414	19.824	1.00 14.76
ATOM	1580	CA	PRO	219	-9.675	-5.138	20.509	1.00 15.37
ATOM	1581	C	PRO	219	-10.433	-4.208	19.567	1.00 19.72
ATOM	1582	0	PRO	219	-9.870	-3.689	18.594	1.00 19.72
ATOM	1583	CB	PRO	219	-8.258	-4.635	20.760	1.00 16.67
ATOM	1584	CG	PRO	219	-7.487	-5.238	19.644	1.00 20.99
ATOM	1585	CD	PRO	219	-8.020	-6.646	19.644	1.00 20.99
	1586	N			-11.700	-3.990	19.892	1.00 10.54
ATOM			MET	220				
MOTA	1587	CA	MET	220	-12.589	-3.173	19.097	1.00 7.76

ATOM	158	18 (ME"	220	-12.67	4 -1.73	8 19.58	6 1.00	9.93
ATOM			ME	r 220	-13.04				12.28
ATOH			B ME	220	-13.98				9.32
ATOM			G MET	220	-14.91	1 -3.270			
ATOM				220	-14.40	9 -3.709			
ATOM			E MET		-14.74	0 -5.479			
MOTA					-12.30	2 -0.813			4.18
MOTA					-12.37	8 0.600			3.81
ATOM					-13.82				8.98
ATOM	159				-14.50		17.929	1.00	7.18
ATOM	159				-11.40			1.00	5.88
ATOM	159				-9.99		18.045	1.00	8.69
ATOM	160				-8.838		17.171	1.00 1	1.36
ATOM	160				-9.319				8.44
ATOM	160				-14.306		19.355	1.00	9.02
MOTA	1603			222	-15.655			1.00 1	0.15
ATOM	1604		LYS	222	-15.826		19.387	1.00 1	2.60
MOTA	1605		LYS	222	-15.204		20.306	1.00 1	2.75
ATOM	1606			222	-16.714		19.823	1.00 1	5.09
ATOM	1607			222	-16.512		21.314	1.00 3	7.80
ATOM				222	-17.741		21.923	1.00 5	0.19
ATOM	1609			222	-17.542		23.389	1.00 6	2.83
ATOM	1610			222	-16.475		23.522	1.00 6	2.90
ATOM	1611		ARG	223	-16.682		18.567	1.00	7.78
ATOM ATOM	1612 1613			223	-16.903	6.122	18.728	1.00	7.43
ATOM	1614	0	ARG	223	-18.216	6.507	18.106	1.00 14	1.04
ATOM	1615	CB		223	-18.471	6.190	16.955	1.00 14	
ATOM	1616	CG	ARG ARG	223	-15.760	6.919	18.065		85
MOTA	1617		ARG	223 223	-16.002	8.432	18.236		. 27
ATOM	1618	NE	ARG	223	-14.808	9.222	17.674	1.00 12	
ATOM			ARG	223	-14.917	9.339	16.230	1.00 21	
ATOM	1620		ARG	223	-15.401 -15.826	10.410	15.670		
ATOM	1621		ARG	223.	-15.459	11.415	16.381	1.00 21	
ATOM	1622	N	LEU	224	-19.051	10.480 7.207	14.373	1.00 36	
ATOM	1623	CA	LEU	224	-20.281	7.729	18.895	1.00 11	
ATOM	1624	C	LEU	224	-19.992	9.123	18.333 17.859	1.00 11	
MOTA	1625	0	LEU	224	-19.508	9.931	18.636	1.00 17	
MOTA	1626	CB	LEU.	224	-21.364	7.768	19.430	1.00 19.	
MOTA	1627	CG	LEU	224	-22.618	8.519		1.00 11.	
MOTA	1628	CD1	LEU	224	-23.329		17.843		
MOTA	1629	CD2	LEU	224	-23.576			1.00 14.	
MOTA	1630	· N	GLY	225	-20.287			1.00 16.	
NOTA	1631	CA	GLY	225	-20.017			1.00 13.	
MOTA	1632	С	GLY	225				1.00 19.	
MOTA	1633	0	GLY	225	-21.026			1.00 22.	
MOTA	1634	N	MET	226	-20.071			1.00 11.	
MOTA	1635	CA	MET	226.	-20.423			1.00 8.	
MOT	1636		MET	226	-19.364			1.00 11.	
MOT	1637	0	MET	226	-18.303			.00 10.	
TOM	1638	CB	MET	226	-20.518			.00 8.	
TOM	1639		MET	226	-21.405			.00 9.	
TOM	1640		MET	226	-23.076			.00 10.	
TOM	1641	CE	MET	226	-23.518			.00 6.3	
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ATO		642	N	PHE	227	-19.	. 665	10.	523 1	1.255	1.00	
ATO	_	643		PHE	227	-18.	691	10.0		0.378		
ATO	_	644	C	PHE	227	-19.				3.931		13.66
ATO	-	645		PHE	227	-20.		10.5		3.580		12.48
ATO		546		PHE	227	-18.	639	8.4		.625	1.00	8.75
ATO		547		PHE	227	-18.	070	8.1		.012		8.71
ATO		548	CD1		227	-16.	694	8.0		.177	1.00	9.74
ATO		49	CD2		227	-18.		8.1		.116	1.00	9.32
ATO			CE1		227	-16.		7.7		.440		11.09
ATO			CE2		227	-18.	402	7.8		.381	1.00	8.85
ATON				HE	227	-17.6		7.6		. 542	1.00	8.29
ATON				SN	228	-17.9		10.14		.086		12.09
ATOM				SN	228	-18.1		10.2		.642		11.99
ATOM				SN	228	-18.5		8.88		.173		19.62
ATOM				SN	228	-17.9		8.34		243	1.00	
MOTA				șn ș	228	-16.8	60	10.75		968	1.00	7.09
MOTA				SN	228	-17.0	41	11.13	_	498	1.00	
ATOH			DD1 A		228	-16.2		11.86	_	946	1.00	
ATOM			D2 AS		228	-18.0	85	10.61		850	1.00	
MOTA				Æ	229	-19.5		8.30		841	1.00 1	
ATOM	166		A II		229	-20.0	01	6.95		565	1.00 1	
ATOM	166				229	-21.2		6.99		687	1.00 1	
ATOM	166	_			229	-22.09		7.90		771	1.00 1	
MOTA	166				229	-20.33	36	6.22		B90	1.00 2	
MOTA :	166		G1 IL		229	-20.76	54	4.780			1.00 2	
	166		G2 IL		229	-21.46	51	6.970			1.00 2	
ATOM	166		D1 IL		229	-19.66	0	3.886	7.1		1.00 3	
ATOM	166				230	-21.37		5.974			1.00 1	
ATOM	167				230	-22.50		5.830			1.00 16	
atom Atom	167		GLI		230	-22.64		4.362	3.5		1.00 20	
ATOM	1672	-	GL		230	-21.79		3.558	3.9		1.00 19	
ATOM	1673 1674				330	-22.33		6.675	2.6		.00 17	
ATOM	1675	_			230	-20.97		6.615	1.9		.00 26	
ATOM	1676				30	-19.96		7.619			.00 46	
ATOM	1677		1 GLN		30	-18.759		7.377	2.5		.00 38	
ATOM	1678		2 GLN		30 .	-20.473		8.748	3.0		.00 44	
ATOM	1679		HIS		31	-23.752		4.010	2.94		.00 19	
ATOM	1680				31	-23.979		2.641	2.53		.00 19	
ATOM	1681	0	HIS		31	-24.244		2.618	1.04		.00 25	
ATOM	1682	СВ	HIS		31	-25.347		2.942	0.59	4 1	.00 25.	. 20
ATOM	1683	CG	HIS		31	-25.158		2.015	3.27	7 1	.00 21.	24
ATOM	1684		HIS HIS		31	-25.361		.571	2.93		.00 25.	66
ATOM	1685	CD2	HIS	23		-25.989	_	.157	1.78		00 28.	29
ATOM	1686		HIS	23		-24.980		.551	3.58		00 29.	45
ATOM	1687		HIS	23		-25.984		.165	1.72		00 29.	07
ATOM	1688	NEZ	CYS	23		-25.376		.617	2.81		00 29.	87
ATOM	1689	CA	CYS	23		-23.230		. 227	0.29	2 1.	00 23.	51
ATOM	1690	C	CYS	23		-23.335		.162	-1.15	1.	00 24.	24
ATOM	1691	0	CYS	23		-24.054			-1.592	1.	00 28.	94
ATOM	1692	CB	CYS	23		-23.603		.216	-1.319	1.	00 27.	88
ATOM	1693	SG	CYS	23		-21.946		190	-1.780	1.	00 25.3	39
ATOM	1694	N	LYS	23:		-21.982			-3.578	1.	00 30.3	39
ATOM	1695	CA	LYS	23		-25.186			-2.260	1.0	00 27.4	4
- .		~~	213	23:	•	-25.913	-0.	094	-2.747	1.6	00 28.8	· 3

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ATO	169	96 C	LYS	233	-25.1	63 -0.79	59 -3.88	34 1.00 35.28
ATO	1 169	7 0	LYS		-25.04			
ATO		98 C	B LYS	233	-27.32		9 3.20	
ATOR			G LYS	233	-28.32			
ATOM		_	D LYS	233	-28.46			5 1.00 58.67
ATON			E LYS		-29.50		8 -0.10	
ATOH			Z LYS	233	-29.59			
ATOH	- •		LYS	234	-24.53			
ATOM			A LYS	234	-23.78	0 -0.48		
ATOM			LYS	234	-22.69			
ATOM		_			-22.53			
MOTA					-23.15	8 0.62°		
ATOM				234	-22.40			8 1.00 49.04
ATOM				234	-22.40	3 1.126		
ATON	1710			234	-22.003		-10.385	1.00 61.46
ATOM	171			234	-22.208		-11:.533	
MOTA	1712			235	-21.951		-4.404	1.00 27.51
MOTA	1713			235	-20.893		-3.904	
ATOH	1714		LEU	235		-2.913	-2.859	
ATOM	1715	-	LEU	235	-20.745		-2.412	1.00 27.78
ATOM	1716	,		235	-19.767		-3.297	1.00 25.05
ATOM	1717		LEU	235	-19.005		-4.226	
ATOM	1718		1 LEU	235	-18.113			1.00 29.02
ATOM	1719 1720		2 LEU	235	-18.198			
ATOM	1720		SER	236			-2.518	
ATOM	1722	CA	SER		-23.363		-1.518	1.00 25.00
ATOM	1723	0	SER	236	-22.546	-3.589	-0.229	1.00 26.77
ATOM	1724	CB	SER SER	236	-22.227	-4.649	0.326	1.00 27.44
ATOM	1725	OG	SER	236 236	-5.555	-5.025	-2.033	1.00 31.15
ATOM	1726	N	SER	237	-24.358	-5.069	-3.188	1.00 47.28
ATOM	1727	CA	SER	237	-22.230	-2.392	0.260	1.00 19.80
ATOM	1728	c c	SER	237	-21.440	-2.266	1.474	1.00 17.87
ATOM	1729	.0	SER	237	-21.371 -21.623	-0.879	2.096	1.00 17.31
ATOH	1730	СВ	SER	237	-20.021	0.124 -2.774	1.440	1.00 15.81
MOTA	1731	OG	SER	237	-19.642	-2.578	1.212	1.00 21.38
MOTA	1732	N	TRP	238	-21.085	-0.842	-0.137	1.00 30.13
MOTA	1733	CA	TRP	238	-20.928	0.407	3.391	1.00 12.22
MOTA	1734	С	TRP	238	-19.531	0.929	4.112 3.774	1.00 11.25
MOTA	1735	0	TRP	238	-18.538	0.193	3.867	1.00 13.06
MOTA	1736	CB	TRP	238	-21.015	0.167	5.612	
MOTA	1737	CG	TRP	238	-22.403	0.081	6.189	1.00 11.29
MOTA	1738	CD1	TRP	238	-23.086	-1.049	6.529	1.00 16.79
MOT	1739	CD2		238	-23.214	1.183	6.607	1.00 13.82
MOT	1740	NE1		238	-24.272	-0.716		1.00 15.81
MOT	1741	CE2		238	-24.382	0.638		1.00 16.81
MOT	1742	CE3		238	-23.076	2.568		1.00 15.44
TOM	1743	CZ2		238	-25.386	1.444		1.00 15.82
TOM	1744	CZ3		238	-24.076	3.364		1.00 16.61
TOM	1745			238,	-25.227	2.799		1.00 16.65
TOM	1746			239	-19.475	2.183		1.00 9.68
TOM				239	-18.231	2.852		1.00 8.63
TOM				239		4.013		1.00 10.72
TOM	1749	0 1	VAL .	239		5.011		1.00 6.54
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MOTA	1750	CB	VAL	239	-18.320	3.369	1.507	1.00 12.66
MOTA	1751	CG1	VAL	239	-17.017	4.042	1.084	
HOTA	1752	CG2	. VAL	239	-18.671	2.225	0.579	1.00 12.01
HOTA	1753	N	LEU	240	-16.871	3.849	4.720	1.00 10.01
ATOM	1754	CA	LEU	240	-16.427	4.799	5.737	1.00 10.33
ATOH	1755	C	LEU	240	-15.128	5.523	5.387	1.00 15.02
MOTA	1756	0	LEU	240	-14.099	4.884	5. 16 6	1.00 14.95
HOTA	1757	CB	LEU	240	-16.229	4.051	7.057	1.00 10.48
MOTA	1758	\sim	LEU	240	-15.548	4.799	8.203	1.00 16.26
ATOM	1759	CD1	LEU	240	-16.472	5.855	8.781	1.00 17.63
HOTA	1760	CD2	LEU	240	-15.124	3.817	9.263	1.00 15.41
MOTA	1761	'n	LEU	241	-15.164	6.850	5.366	1.00 12.02
MOTA	1762	CA	LEU	241	^{-13.962}	7.620	5.070	1.00 12.27
MOTA	1763	С	LEU	241	-13.329	8.283	6.298	1.00 18.80
MOTA	1764	0	LEU	241	-13.777	9.334	6.774	1.00 18.43
HOTA	1765	CB	LEU	241	-14.214	8.648	3.964	1.00 11.42
HOTA	1766	CG	LEU	241	-14.176	8.089	2.541	1.00 15.05
HOTA	1767	CD1	LEU	241	-15.379	7.211	2.276	1.00 15.36
ATOM	1768		LEU	241	-14.119	9.229	1., 547	1.00 16.74
ATOM	1769	N	MET	242	-12.284	7.641	6.806	1.00 17.21
ATOM	1770	CA	MET	242	-11.544	8.120	7.967	1.00 18.01
ATOM	1771	С	MET	- 242	-10.409	9.053	7.550	1.00 23.47
ATOM	1772	0	MET	242	-9.468	8.668	6.864	1.00 24.19
MOTA	1773	СВ	MET	242	-10.966	6.950	8.766	1.00 20.26
ATOM	1774	CG	MET	242	-11.988	6.085	9.455	1.00 22.94
ATOM	1775	SD	MET	242	-12.629	6.786	10.951	1.00 25.83
ATOM	1776	CE	MET	242	-11.268	6.581	12.006	1.00 22.11
ATOM	1777	N	LYS	243	-10.562	10.302	8.032	1.00 18.99
ATOM	1778	CA	LYS	243	-9.548	11.312	7.802	1.00 17.66
ATOM	1779	С	LYS	243	-8.317	11.048	8.622	1.00 17.67
ATOM	1780	0	LYS	243	-8.398	10.740	9.802	1.00 18.08
ATOM	1781	CB	LYS	243	-10.105	12.714	8.132	1.00 21.90
ATOM	1782	CG ·	LYS	243	-10.690	12.755	9.559	1.00 25.80
MOTA	1783	CD	LYS	243	-12.103	12.137	9.582	1.00 18.62
MOTA	1784	CE	LYS	243	-12.461	11.654	11.002	1.00 26.35
ATOM	1785	NZ	LYS	243	-13.822	11.096	11.020	1.00 23.36
MOTA	1786	N	TYR	244	-7.159	11.177	7.954	1.00 13.01
ATOM	1787	CA	TYR	244	-5.890	10.998	8.667	1.00 12.02
ATOM	1788	C	TYR	244	-5.195	12.310	8.951	1.00 19.96
ATOM	1789	0	TYR	244	-5.138	13.207	8.100	1.00 20.97
ATOM	1790	СВ	TYR	244	-4.898	10.149	7.870	1.00 11.09
ATOH	1791	CG	TYR	244	-4.867	8.706	8.243	1.00 12.20
ATOM	1792	CD1	TYR	244	-4.636	8.316	9.553	1,00 12.28
ATOM	1793	CD2		244	-5.089	7.722	7.285	1.00 14.62
MOTA	1794	CE1		244	-4.637	6.974	9.901	1.00 13.09
MOTA	1795		TYR	244	-5.089	6.376	7.624	1.00 12.96
MOTA	1796	CZ	TYR	244	-4.864	6.016	8.927	1.00 23.64
ATOM	1797	ОН	TYR	244	-4.858	4.692	9.255	1.00 34.09
MOTA		N	LEU	245	-4.618	12.389	10.141	1.00 18.61
ATOM	1799	CA	LEU	245	-3.873	13.559	10.553	1.00 19.17
MOTA	1800	C	LEU	245	2.578	13.463	9.768	1.00 28.06
ATOM	1801	ō	LEU	245	-1.675	12.710	10.122	1.00 29.74
MOTA	1802	СВ	LEU	245	-3.586	13.518	12.056	1.00 18.84
ATOM	1803	CG	LEU	245	-2.913	14.754	12.632	1.00 22.04
								

ATOM	1804	CI	1 LEU	245	-3.868	15.926	12.519	1.00 21.70
MOTA	1809	C	2 LEU	245	-2.503	14.504	14.070	
MOTA	1806	N	GLY	246	-2.544	14.157	8.644	
ATOM	1807	CA	GLY	246	-1.381	14.155	7.782	1.00 28.42
MOTA	1808		GLY	246	-1.875	14.635	6.435	1.00 34.25.
ATOM	1809		GLY	246	-2.143	15.817	6.252	1.00 38.08
ATOM	1810		ASN	247	-2.098		5.517	1.00 26.63
ATOM	1811			247	-2.575	14.073	4.198	
ATOM	1812		asn	247	-3.077	12.827	3.491	
MOTA	1813		asn	247	-3.054	12.741	2.262	
ATOM	1814			247	-1.458		3.388	
MOTA	1815			247	-0.157	13.956	3.414	
ATOM	1816		1 ASN	247	0.368	13.645	4.487	
ATOM	1817		2 ASN	247	0.373	13.637	2.237	
MOTA	1818		ALA	248	-3.538	11.863	4.276	
ATOM	1819	-		248	-4.063	10.619	3.735	
atom	1820 1821	C	ALA	248	-5.469	10.366	4.300	1.00 17.56
ATOM	1822		ALA	248	-5.809	10.858		1.00 14.85
ATOM	1823	CB N	ALA THR	248	-3.125	9.458	4.064	
ATOM	1824	CA	THR	249 249	-6.286 -7.651	9.648	3.535	1.00 11.98
ATOM	1825	C	THR	249	-7.837	9.314 7.808	3.912 3.810	1.00 9.23
ATOM	1826	0	THR	249	-7.495	7.187	2.808	1.00 11.01 1.00 9.71
ATOM	1827	СВ	THR	249	-8.689		2.996	1.00 5.40
ATOM	1828	OG:		249	-8.623	11.435	3.175	1.00 6.92
MOTA	1829	CG2		249	-10.098	9.525	3.302	1.00 1.00
ATOM	1830	N	ALA	250	-8.331	7.233	4.891	1.00 7.89
MOTA	1831	CA.		250	-8.601	5.819	5.001	1.00 7.37
ATOM	1832	С	ÁLA	250	-10.008	5.556	4.497	1.00 11.92
ATOM	1833	0	ALA	250	-10.950	6.222	4.927	1.00 14.56
MOTA	1834	CB	ALA	250	-8.504	5.396	6.468	1.00 7.88
ATOM	1835	N	ILE	251	-10.146	4.646	3.542	1.00 5.32
MOTA	1836	CA	ILE	251	-11.457	4.273	3.030	1.00 3.97
MOTA	1837	С	ILE	251	-11.611	2.810	3.441	1.00 6.37
MOTA	1838	0	ILE	251	-10.709	1.998	3.225	1.00 6.05
MOTA	1839	CB	ILE	251	-11.564	4.365	1.483	1.00 7.21
ATOM	1840	CG1		251	-11.090	5.725	0.970	1.00 6.26
ATOM	1841	CG2		251	-13.007	4.160	1.052	1.00 10.19
ATOM	1842	CD1		251	-11.354	5.922	-0.507	1.00 1.00
MOTA	1843	N	PHE	252	-12.714	2.501	4.112	1.00 1.97
MOTA	1844	CA	PHE	252	-13.006	1.147	4.572	1.00 1.00
ATOM ATOM	1845 1846	C	PHE	252	-14.298	0.701	3.928	1.00 7.08
ATOM	1847	. 0	PHE	252	-15.267	1.458	3.893	1.00 8.36
ATOM	1848	CB CG	PHE	252	-13.196 -11.989	1.112	6.094	1.00 1.68
ATOM	1849		PHE PHE	252 252	-11.989	1.534	6.877	1.00 1.01
ATOM	1850		PHE	252 252	-11.936	0.676	7.019	1.00 2.49
ATOM	1851		PHE	252 ·	-11.936 -9.792	2.785	7.466	1.00 1.00
ATOM	1852	CE2	PHE	252	-10.827	1.052	7.745	1.00 2.81
ATOM	1853	CZ	PHE	252 252		3.169 2.302	8.192 8.328	1.00 4.34
ATOM	1854	N	PHE	252		-0.556	3.507	1.00 2.20
ATOM	1855	CA	PHE	253 253	the second secon	-1.110	2.858	1.00 3.93 · 1.00 4.58
ATOM	1856	C	PHE	253		-2.354		1.00 4.58
ATOM	1857	0	PHE	253		-3.388		1.00 13.67
		-				3.330	J. J. L	11.47

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ATOH	185	8 C	в Рив	253	-15.17	4 -1.50	0 1.41	1.00	5.96
ATON	185	9 (1	G PHE	253	-14.64	1 -0.37	7 0.57		7.08
ATOM	186	0 CI	D1 PHE	253	-15.50	9 0.479			8.18
ATOM	186	1 CI	D2 PHE	253	-13.27	4 -0.162	0.469		10.35
ATOM	. 186	2 CI	El PHE	253	-15.02	1 1.526	-0.863		10.14
ATOM	186	3 CI	E2 PHE	253	-12.78	1 0.886	-0.301		10.64
ATOM	1864		PHE		-13.65		-0.965	1.00	8.11
ATOM	1869		LEU	254	-17.14	2 -2.265	4.212	1.00	15.28
ATOM	1866			254	-17.73	6 -3.383	4.954	. 1.00	15.29
ATOM	1867		LEU	254	-18.79			1.00	19.45
MOTA	1868		LEU	254	-19.90	8 -3.482	3.949	1.00	20.52
MOTA	1869			254	-18.372		6.259		14.77
MOTA				254	-18.94		7.192	1.00	18.06
MOTA	1871		1 LEU	254	-17.83		7.607	1.00	18.39
ATOM	1872			254	-19.564			1.00	17.27
MOTA	1873		PRO	255	-18.459			1.00	14.39
MOTA	1874			255	-19.415			1.00	
MOTA			PRO	255	-20.653			1.00	22.98
MOTA	1876		PRO	255	-20.571		4.370	1.00	
MOTA	1877		PRO	255	-18.596		1.905	1.00	
MOTA	1878		PRO		-17.692		3.041	1.00 1	
MOTA	1879		PRO	255	-17.225			1.00 1	
MOTA	1880	N ·		256	-21.794		2.532	1.00 1	
ATOM	1881	CA	ASP	256	-23.057		3.108	1.00 1	
ATOM	1882 1883	c O	ASP	256	-22.960		3.426	1.00 2	
ATOM	1884	CB	asp asp	256 256	-21.892		3.325	1.00 2	
ATOM	1885	CG	ASP	256 256	-24.249		2.169	1.00 1	
ATOM	1886		ASP	256	-24.732 -24.129	-4.970 -4.149	2.262	1.00.2	
ATOM	1887		ASP	256	-25.730	-4.635	2.985	1.00 2	
ATOM	1888	N	GLU	257	-24.057	-8.818	1.570 3.893	1.00 3	
ATOM	1889	CA	GLU	257		-10.240		1.00 2	
MOTA	1890	c c	GLU	257			2.909	1.00 2	
ATOM	1891	0	GLU	257		-10.862	1.953	1.00 2	
ATOM	1892	CB	GLU	257		-10.629	4.756	1.00 2	
MOTA	1893	CG	GLU	257	-25.538		5.444	1.00 4	
ATOM	1894	CD	GLU	257	-26.943		5.968	1.00 72	
MOTA	1895	OE1	GLU	257	-27.634		6.418	1.00 76	
MOTA	1896	OE2		257	-27.352		5.927	1.00 67	
MOTA	1897	N	GLY	25B	-22.613		2.839	1.00 23	
MOTA	1898	CA	GLY	258	-22.228		1.648	1.00 22	
MOTA	1899	С	GLY	258	-21.875	-11.549	0.407	1.00 28	
MOTA	1900	0	GLY	258	-21.772	-12.107	-0.685	1.00 30	
MOTA	1901	N	LYS	259	-21.645		0.561	1.00 21	
MOTA	1902	CA	LYS	259	-21.319	-9.397	-0.581	1.00 19	
MOTA	1903	С	LYS	259	-19.854	-8.959	-0.663	1.00 21	
MOTA	1904	0	LYS	259	-19.564	-7.857	-1.134	1.00 21	
MOTA	1905	CB	LYS	259	-22.233	-8.171	-0.595	1.00 21	
MOTA	1906	CG	LYS	259	-23.669	-8.469	-0.980	1.00 25	
MOTA	1907	CD	LYS	259	-23.822	-8.619		1.00 29	
MOT	1908	CE	LYS	259	-25.263		-2.885	1.00 37	. 09
MOT	1909	NZ	LYS	259	-25.394			1.00 46	
TOM	1910	N	LEU	260	-18.924		-0.262	1.00 16	. 31
MOT	1911	CA	LEU	260	-17.500	-9.474	-0.322	1.00-14	.94

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MOTA	1912	C	LEU	260	-16.929	-9.593	-1.732	1.00 26.00
ATOH .	1913	0	LEU	260	-16.263	-8.677	-2.213	1.00 27.32
ATOM	1914	·CB	LEU	260	-16.667		0.672	1.00 12.73
ATOM	1915	CG	LEU	260	-15.138	-10.131	0.721	1.00 13.73
HOTA	1916	CD1	LEU	260	-14.696	-8.701	0.938	1.00 11.63
ATOM	1917	CD2	LEU	260	-14.611	-11.018	1.820	1.00 16.86
ATOM	1918	N	GLN	261		-10.688	-2.414	1.00 25.25
ATOM	1919	CA	GLN	261	-16.743	-10.894	-3.775	1.00 27.07
ATOM	1920	C	GLN	261	-17.251	-9.774	-4.693	1.00 30.23
ATOM	1921.	ō	GLN	261 .	-16.502	-9.238	-5.508	1.00 32.36
ATOM	1922	CB	GLN	261	-17.181	-12.266	-4.303	1.00 29.06
ATOM	1923	CG	GLN	261	-16.633	-12.612	-5.683	1.00 46.56
ATOM	1924	CD	GLN	261	-15.127	-12.692	-5.719	1.00 84.32
MOTA	1925		GLN	261		-12.444	-4.724	1.00 84.95
ATOM	1926	NE2	GLN	261	-14.592	-13.051	-6.873	1.00 87.30
ATOM	1927	N	HIS	262	-18.509	-9.392	-4.495	1.00 21.14
ATOM	1928	CA	HIS	262	-19.147	-8.333	-5.259	1.00 18.96
ATOM	1929	C	HIS	262	-18.394	-7.009	-5.079	1.00 20.50
ATOM	1930	ō	HIS	262	-18.037	-6.354	-6.052	1.00 19.00
	1931	СВ	HIS	262	-20.600	-8.177	-4.791	1.00 19.08
ATOM	1932	CG	HIS	262	-21.325	-7.025	-5.418	1.00 21.74
ATOM	1933		HIS	262	-22.215	-7.191	-6.455	1.00 23.34
ATOM	1934		HIS	262	-21.298	-5.701	-5.154	1.00 23.53
ATOM	1935		HIS	262	-22.707	-6.018	-6.810	1.00 22.73
ATOM	1936		HIS	262	-22.163	-5.096	-6.034	1.00 23.15
ATOM	1937	N	LEU	263	-18.170	-6.622	-3.829	1.00 15.36
ATOM	1938	CA	LEU	263	-17.460	-5.391	-3.533	1.00 14.39
ATOM	1939	c	LEU	263	-16.122	-5.351	-4.261	1.00 18.62
ATOM	1940	0	LEU	263	-15.841	-4.423	-5.023	1.00 17.02
ATOM	1941	СВ	LEU	263	-17.233	-5.281	-2.027	1.00 14.57
ATOM	1942	CG	LEU	263	-16.264	-4.205	-1.525	1.00 19.15
ATOM	1943		LEU	263	-16.837	-2.811	-1.731	1.00 18.55
ATOM	1944		LEU	263	-15.965	-4.450	-0.058	1.00 22.85
ATOM	1945	N	GLU	264	-15.344	-6.412	-4.083	1.00 17.48
MOTA	1946	CA	GLU	264	-14.020	-6.530	-4.686	1.00 17.94
ATOM	1947	C	GLU	264	-14.049	-6.405	-6.205	1.00 23.91
ATOM	1948	0	GLU	264	-13.098	-5.912	-6.819	1.00 23.40
MOTA	1949	СВ	GLU	264	-13.399	-7.868	-4.306	1.00 18.87
ATOM	1950	CG	GLU	264	-13.258	-8.104	-2.833	1.00 19.89
ATOM	1951	CD	GLU	264	-12.550	-9.399	-2.532	1.00 37.72
ATOM	1952	OE1	GLU	264		-10.410	-3.201	1.00 44.58
ATOM	1953		GLU	264	-11.721	-9.410	-1.603	1.00 30.00
MOTA	1954	N	ASN	265	-15.137	-6.882	-6.797	1.00 22.09
MOTA	1955	CA	ASN	265	-15.314	-6.849	-8.239	1.00 22.25
ATOM	1956	C ·	ASN	265	-15.950	-5.561	-8.753	1.00 27.73
ATOM	1957	0	ASN	265	-15.660	-5.140	-9.869	1.00 29.41
ATOM	1958	СВ	ASN	265	-16.130	-8.060	-8.687	1.00 22.20
MOTA	1959	CG	ASN	265	-15.444	-9.370	-8.354	1.00 53.15
ATOM	1960	OD1	ASN .	265	-14.220	-9.492	-8.455	1.00 57.34
ATOM	1961		ASN	265	-16.222	-10.355	-7.945	1.00 43.73
ATOM	1962	N	GLU	266	-16.765	-4.909	-7.925	1.00 23.88
ATOM	1963	CA	GLU	266	-17.443	-3.671	-8.319	1.00 22.90
ATOM	1964	c i	GLU	266	-16.629	-2.405	-8.094	1.00 22.66
ATOM	1965	0	GLU	. 266	-16.998	-1.341	-8.582	1.00 22.09

							
ATOM	1966	CB	GLU	266	-18.799	-3.528 -7.6	
ATOM	1967	CC	GLU	266	-19.788	-4.625 -7.9	-
MOTA	1968	CD	GLU	266	-19.966	-4.898 -9.3	
MOTA	1969	OE1	GLU	266	-19.961	-3.949 -10.1	
MOTA	1970	OE2	GLU	266	-20.118	-6.089 -9.7	
MOTA	1971	N	LEU	267	-15.540	-2.510 -7.3	
MOTA	1972	CA	LEU	267	-14.677	-1.363 -7.0	•
HOTA	1973	C	LEU	267	-14.142	-0.753 -8.3	
MOTA	1974	0	LEU	267	-13.696	-1.476 -9.2	
ATOM	1975	CB	LEU	267	-13.502	-1.792 -6.1	
MOTA	1976	CG	LEU	267	-13.838	-2.150 -4.7	
MOTA	1977	CD1	LEU	267	-12.623	-2.724 -4.0	
ATOM	1978	CD2	LEU	267	-14.367	-0.924 -3.9	
ATOM	1979	N.	THR	268	-14.178	0.574 -8.4	•
HOTA	1980	CA	THR	268	-13.693	1.278 -9.6	
ATOM	1981	С	THR	268	-13.074	2.610 -9.2	
ATOM	1982	0	THR	268	-13.603	3.273 -8.3	
	1983	CB	THR	268	-14.849	1.596 -10.6	
ATOM	1984	OG1	THR	268	-15.647	2.688 -10.1	50 1.00 29.63
ATOM	1985	CG2	THR	268	-15.740	0.391 -10.8	64 1.00 23.81
ATOH	1986	N	HIS	269	-11.998	3.031 -9.8	76 1.00 16.16
ATOM	1987	CA	HIS .	269	-11.353	· 4.297 -9.5	25 1.00 17.99
ATOM	1988	С	HIS	269	-12.393	5.422 -9.40	34 1.00 28.56
ATOM	1989	0	HIS	269	-12.331	6.298 -8.63	17 1.00 29.22
ATOM	1990	CB	HIS	269	-10.222	4.639 -10.5	12 1.00 18.15
ATOM	1991	CG	HIS	269	-9.671	6.032 -10.3	9 1.00 21.26
ATOM	1992		HIS	269	-9.487	6.873 -11.43	37 1.00 23.32
ATOM	1993		HIS	269	-9.255	6.718 -9.2	73 1.00 23.72
ATOM	1994		HIS	269	-8.978	8.019 -11.03	
ATOM	1995		HIS	269	-8.827	7.954 -9.71	12 1.00 23.46
ATOM	1996	N	ASP	270	-13.376	5.334 -10.37	78 1.00 27.36
ATOM	1997	CA	ASP	270	-14.452	6.318 -10.47	71 1.00 28.28
	1998	C.	ASP	270	-15.204	6.391 -9.14	
ATOM	1999	0	ASP	270	-15.198	7.424 -8.48	34 1.00 28.48
ATOM	2000	CB	ASP	270	-15.415	5.934 -11.61	
ATOM	2001	CG	ASP	.270	-16.509	6.984 -11.87	
ATOM	2002	ODI	ASP	270	-16.777	7.870 -11.03	1.00 60.70
MOTA	2003		ASP	270	-17.125	6.907 -12.96	
ATOM	2004	N ·	ILE	271	-15.783	5.263 -8.74	18 1.00 23.52
ATOM	2005	CA	ILE	271	-16.561	5.147 -7.51	
ATOM	2006	С	ILE	271	-15.771	5.628 -6.30	0 1.00 22.26
MOTA	2007	0	ILE	271	-16.306	6.309 -5.42	
ATOM	2008	CB	ILE	271	-17.015	3.682 -7.27	9 1.00 26.80
MOTA	2009	CG1	ILE	271	-17.922	3.220 -8.41	9 1.00 28.03
ATOM	2010		ILE	271	-17.769	3.546 -5.98	6 1.00 28.26
ATOM	2011	CD1	ILE	271	-18.340	1.771 -8.31	.5 1.00 35.59
ATOM	2012	N.	ILE	272	-14.479	5.327 -6.28	
ATOM	2013	CA	ILE	272	-13.629	5.723 -5.16	9 1.00 13.20
ATOM	2014	C	ILE	272.	-13.386	7.220 -5.19	4 1.00 19.00
ATOM	2015	ō	ILE	272	-13.359	7.867 -4.14	4 1.00 19.72
ATOM	2016	СВ	ILE	272	-12.333	4.890 -5:12	7 1.00 14.50
ATOM	2017		ILE	272	-12.697	3.447 -4.74	0 1.00 13.26
ATOM	2018		ILE	272	-11.356	5.477 -4.12	7 1.00 14.10
ATOM	2019		ILE	272	-11.552	2.478 -4.76	1. 1.00 24.26

								•
ATO	1 202	20 1	TH:	R 273	-13.29	3 7.78	5 -6.391	1.00 18.09
ATOR	202	21 (CA THE					
ATOP	9 202	22 (-14.40			
ATOM			THE					
ATON	202	24 C	В ТН		-12.76			
ATOM		-	G1 THE	R ' 273	-11.56			
ATOM			G2 THE		-12.55			
ATOM		7 N	LYS	274	-15.51			
ATOM		_	A LYS	274	-16.83			
MOTA			LYS	274	-16.87			
ATOM			LYS	274	-17.29			
ATOM				274	-17.880		•	1.00 26.15
ATOM					-19.293	9.228		1.00 50.14
MOTA				274	-20.187		-7.193	1.00 66.22
ATOM	2034				-19.701	7.754	-8.580	1.00 82.49
ATOM					-20.447	6.600	-9.135	1.00 95.34
ATOH	2036				-16.370			1.00 23.58
ATOM	2037				-16.346		-2.306	1.00 21.54
ATOM			PHE		-15.431			1.00 21.45
ATOM	2039			275	-15.793		-0.898	1.00 21.56
ATOM .				275	-15.907		-1.749	1.00 23.59
ATOM	2041			. 275	-16.911		-1.974	1.00 24.46
ATOM	2042		1 PHE	275	-18.236		-1.561	1.00 26.80
ATOM	2043		2 PHE	275		5.197	-2.609	1.00 26.12
ATOM	2045		2 PHE	275	-19.176	5.523	-1.778	1.00 26.69
ATOM	2046			275 275	-17.488	4.196	-2.828	1.00 29.22
ATOM	2047	N	LEU	275	-18.805 -14.258	4.364	-2.412	1.00 26.65
MOTA	2048	CA		276	-13.300	10.062	-2.414	1.00 16.44
MOTA	2049	С	LEU	276	-13.894	11.086 12.484	-2.000	1.00 15.37
MOTA	2050	0.	LEU	276	-13.819	13.255	-2.078	1.00 24.02
ATOM .	2051	СВ	LEU	276	-12.030	11.019	-1.123 -2.839	1.00 25.62
ATOM -	2052	CG	LEU	276	-11.187	9.776	-2.553	1.00 19.14
ATOM	2053	CD	LEU.	276	-9.889	9.868	-3.336	1.00 19.14
MOTA	2054	CD2	LEU	276	-10.887	9.649	-1.054	1.00 22.14
MOTA	2055	N	GLU	277	-14.539	12.781	-3.200	1.00 22.07
ATOM	2056	CA	GLU	277	-15.162	14.077		1.00 22.27
MOTA	2057	C	GLU	277	-16.323	14.256		1.00 30.33
MOTA	2058	0	GLU	277	-16.852	15.359		1.00 34.24
MOTA	2059	CB	GLU	277	-15.651	14.231		1.00 23.70
ATOM	2060	CG	GLU	277	-14:592	13.986		1.00 37.77
ATOM	2061	CD	GLU	277	-13.401	14.944		1.00 65.64
atom Atom	2062 2063		GLU	277	-13.273			1.00 72.39
ATOM .	2064		GLU	277	-12.577		-6.836	1.00 57.27
ATOM	2065	N	ASN	278	-16.705			1.00 24.83
ATOM	2066	CA	ASN	278	-17.788			1.00 23.02
ATOM	2067	С О	ASN	278	-17.337	13.798		1.00 23.90
ATOM	2068	CB	ASN	278	-16.453	13.251		00 20.54
ATOM	2069	တ	asn Asn	278 278				00 24.64
MOTA	2070	OD1		278 278		11.899		.00 63.15
NTOM	2071	ND2		278 278		10.950		.00 59.86
MOT	2072	N N	GLU .	278 . 279		13.018		.00 61.71
TOM	2073	CA	GLU	279		14.903		.00 23.66
			7 20	213	-17.664	15.588	2.192 1	.00 24.15

							•	
MOTA	2074	C	GLU	279	-18.810	15.596	3.209	1.00 28.94
MOTA	2075	0	GLU	279	-18.790	16.360	4.163	1.00 29.34
MOTA	2076	CB	GLU	279	-17.185	17.004	1.905	1.00 25.78
MOTA	2077	CC	GLU	279	-15.827	17.082	1.214	1.00 44.13
ATOM	2078	СĎ	GLU	279	-15.420	18.506	0.860	1.00 92.58
MOTA	2079	OE	GLU	279	-15.927	19.466	1.482	1.00100.65
MOTA	2080	0E2	GLU	279	-14.574	18.664	-0.042	1.00100.57
MOTA	2081	N	ASP	280	-19.791	14.717	3.016	
MOTA	2082	CX	ASP	280	-20.933	14.607	3.920	1.00 22.02
MOTA	2083	C	ASP	280	-20.564	13.769	5.137	1.00 22.41
MOTA	2084	0	ASP	- 280	-19.554	13.071	5.138	1.00 22.95
MOTA	2085	CB	ASP	280	-22.108	13.957	3.209	1.00 24.35
MOTA	2086	CC	ASP	280	-22.693	14.833	2.151	1.00 42.62
MOTA	2087	OD1	ASP	280	-23.039	15.985		1.00 45.44
MOTA	2088		ASP	280	-22.818	14.362	0.998	1.00 52.57
MOTA	2089	N	ARG	281	-21.420	13.871	6.172	1.00 14.63
ATOM	2090	CX	ARG	281	-21.201	13.094	7.380	1.00 11.86
MOTA	2091	C	ARG	281	-22.517	12.866	8.067	1.00 15.11
ATOM	2092	0	ARG	281	-23.465	13.601	7.840	1.00 15.94
MOTA	2093	CB	ARG	281	-20.312	13.857	8.381	1.00 10.33
MOTA	2094	CG	ARG	281	-18.855	13.944		1.00 19.95
ATOM	2095	CD.	ARG	281 1	-18.034	14.659	8.979	1.00 9.99
MOTA	2096	NE	ARG	281	-16.672	14.863	8.523	1.00 12.05
MOTA	2097	CZ	ARG	281	-15.699	14.096	8.926	1.00 20.69
MOTA	2098	NH1	ARG	281	-15.913	13.105	9.741	1.00 1.00
MOTA	2099	NH2	ARG	281	-14.491	14.329	8.503	1.00 29.15
MOTA	2100	N	ARG	282	-22.567	11.832	8.927	1.00 10.27
MOTA	2101	CX	ARG	282	-23.782	11.605	9.690	1.00 9.47
ATOM	2102	С	ARG	282	-23.459	10.913	10.984	1.00 16.55
ATOM	2103	0	ARG	~282	-22.391	10.338	11.117	1.00 17.04
MOTA	2104	CB	ARG		-24.841	10.828	8.881	1.00 3.07
MOTA	2105	CG	ARG	282	-24.354	9.403	8.551	1.00 5.84
MOTA	2106	CD	ARG	282	-25.536	8.601	7.978	1.00 25.67
ATOM	2107	NE	ARG	282	-25.077	7.326	7.458	1.00 38.49
MOTA	2108	CZ	ARG	282	-25.151	7.044	6.188	1.00 48.45
MOTA	2109	NH1	ARG	282	-25.622	7.902	5.330	1.00 26.72
MOTA	2110	NH2	ARG	282	-24.745	5.883	5.771	1.00 34.81
MOTA	2111	N	SER	283	-24.392	10.980	11.953	1.00 12.72
ATOM	2112	CA	SER	283	-24.123	10.333	13.224	1.00 11.68
MOTA	2113	С	SER	283	-24.128	8.839	13.064	1.00 12.67 1.00 12.85
MOTA	2114	0	SER	283	-24.929	8.308	12.313 14.317	
MOTA	2115	CB	SER	283	-25.111	10.784		1.00 17.91
ATOM	2116	OG	SER	283	-26.441	10.375	13.989	1.00 28.36 1.00 8.25
ATOM	2117	N	ALA	284	-23.205	8.167	13.778	
MOTA	2118	CA	ALA	284	-23.139	6.721	13.670	1.00 7.48
MOTA	2119	C	ALA	284	-22.279	6.151	14.763 15.319	1.00 10.55 1.00 7.46
ATOM .	2120	0	ALA	284	-21.442 -22.601	6.846	12.293	1.00 7.46
ATOM	2121	CB	ALA	284	-22.513	6.288 4.860	15.071	1.00 8.44
MOTA	2122	N	SER	285	-22.513 -21.746	4.231	16.131	1.00 10.17
MOTA	2123	CA	SER	285 285	-20.691	3.352	15.519	1.00 16.06
ATOM	2124 2125	С 0	SER	285 285	-20.897	2.161		1.00 19.28
MOTA MOTA	2125	CB	SER SER	285 285	-22.719	3.436	17.021	1.00 17.06
ATOM	2126	OG	SER	285 285	-23.635	4.344	17.641	1.00 34.64
WI ON	2121	W.	254	463	-23.033	4.344	* /	

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ATOH	2128	N	LEU	286	-19.539	3.969	15.197	1.00 8.68
ATON	2129	ĊA	LEU	286	-18.495	3.216	14.531	1.00 7.51
HOTA	2130	C	LEU	286	-17.713	2.313	15.464	1.00 13.65
MOTA	2131	0	LEU	286	-17.269	2.749	16.535	1.00 14.46
HOTA	2132	CB	LEU	286	-17.510	4.166	13.833	1.00 7.11
MOTA	2133	CG	LEU	286	-16.310	3.517	13.130	1.00 10.35
MOTA	2134	CD:	1 LEU	286	-16.783	2.793	11.883	1.00 10.07
ATOH	2135	CD	2 LEU	286	-15.277	4.552	12.771	1.00 10.77
MOTA	2136	N	HIS	287	-17.564	1.052	15.060	1.00 10.80
ATOM	2137	CA	HIS	287	-16.767	0.068	15.801	1.00 12.55
ATOM	2138	С	HIS	287	-15.702	-0.357	14.803	1.00 13.43
MOTA	2139	0	HIS	287	-15.989	-1.099	13.871	1.00 14.75
MOTA	2140	CB	HIS	287	-17.580	-1.170	16.170	
ATOM	2141	CC	HIS	287	-18.581	-0.946	17.257	1.00 19.45
MOTA	2142		HIS	287	-19.745	-0.235	17.064	
MOTA	2143		HIS	287	-18.618	-1.394	18.531	1.00 21.47
MOTA	2144		HIS	287	-20.462	-0.253	18.172	1.00 21.49
ATOM	2145		HIS		-19.796	•		1.00 21.79
MOTA	- 2146	N	LEU	288	-14.479		15.000	1.00 4.50
MOTA	2147	CA	LEU	288	-13.391	-0.230	•	1.00 2.17
ATOM	2148	С	LEU	288	-12.321	-0.947	14.904	1.00 11.97
ATOH	2149	0	LEU	288	-11.891	-0.449	15.943	1.00 12.25
MOTA	2150	CB	LEU	288	-12.841	1.046	13.467	1.00 1.00
MOTA	2151	CG	LEU	288	-11.665	0.921	12.504	1.00 3.34
ATOM	2152		LEU	288	-12.073	0.170	11.231	1.00 2.56
ATOM	2153		LEU	288	-11.163	2.312		1.00 2.67
ATOM	2154 2155	N .		289 - 289	-11.911 -10.883	-2.150 -2.892	14.469 15.200	1.00 13.97
ATOM	2156	CA			-9.483	-2.832	15.154	1.00 22.55
ATOM	2157	0	PRO PRO	289 289	-9.046	-1.715	14.140	1.00 22.35
ATOM	2158	CB	PRO	289	-10.889	-4.257	14.506	1.00 17.38
ATOM	2159	CG	PRO	289	-11.304	-3.925	13.106	1.00 21.04
ATOM	2160	CD	PRO	289	-12.422	-2.953	13.338	1.00 15.79
ATOM	2161	N	LYS	290	-8.779	-2.419	16.262	1.00 19.44
ATOM	2162	CA	LYS	290	-7.421	-1.945	16.392	1.00 18.93
MOTA	2163	C	LYS	290	-6.631	-3.126	15.838	1.00 24.83
ATOM	2164	ō	LYS	290	-6.953	-4.279	16.151	1.00 25.89
ATOM	2165	CB	LYS	290	-7.123	-1.718	17.874	1.00 20.28
ATOM	2166	CG	LYS	290	-5.749	-1.192	18.216	1.00 35.38
MOTA	2167	CD	LYS	290	-5:683	-0.954	•	1.00 53.02
MOTA	2168	CE.	LYS	290	-4.319	-0.512	20.181	1.00 77.66
MOTA	2169	NZ	LYS	290	-4.355	-0.183	21.634	1.00 94.71
ATOM	2170	N	LEU	291	-5.663	-2.859	14.963	1.00 22.36
MOTA	2171	CA	LEU	291 -	-4.862	-3.928	14.375	1.00 22.32
ATOM	2172	С	LEU	291	-3.587	-3.451	13.708	1.00 25.74
MOTA	2173	0	LEU	291	-3.507	-2.315	13.271	1.00 26.04
HOTA	2174	CB	LEU	291	-5.695	-4.711	13.356	1.00 22.24
MOTA	2175	CG	LEU	291	-5.983	-4.114	11.974	1.00 26.16
MOTA	2176		LEU	291	-6.823	-5.127	11.218	1.00 26.63
MOTA	2177		LEU	291	-6.688	-2.778	12.043	1.00 28.41
MOTA	2178	N	SER	292	-2.600	-4.335	13.619	1.00 21.54
MOTA	2179	CA	SER	292	-1.343	-4.005	12.973	1.00 22.64
MOTA	2180	C	SER	292	-1.020		11.999	1.00 29.52
ATOM	2181	0	SER	292	-0.215	-6.017	12.310	1.00 31.69

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MOTA	2182	CB	SER	292	-0.242	-3.869	14.011	1.00 27.63	
ATOM	2183	œ	SER	292	-0.23				
ATOM	2184	N	ILE	293	-1.655	-5.102	10.827	1.00 23.99	
MOTA	2185	CA	ILE	293	-1.450			1.00 21.90	
ATOM	2186	C	ILE	293	-0.240		8.939		
ATOM	2187	0	ILE		0.239				
atom	2188	CB	ILE	293	-2.721				
MOTA	2189		1 ILE	293	-3.077		8.167		
MOTA	2190		2 ILE	293	-3.881		9.851		
MOTA	2191	CD:		293	-4.156		7.138		
MOTA	2192	N	THR	294	0.207				
ATOM	2193	CA	THR	294	1.360		7.416		
MOTA	2194	С	THR	294	1.078		6.098		
ATOM	2195	0	THR	294	0.112		5.977		
ATOM	2196	CB	THR	294	2.572		8.089		
MOTA	2197		I THR	294	2.622	-7.140	9.462	1.00 35.90	
MOTA	2198	CG		294	3.857	-7.128	7.405	1.00 30.05	
ATOM	2199	N	GLY	295	1.900	-7.312	5.099	1.00 16.51	
MOTA	2200 2201	CA C	GLY	295	1.749		3.805	1.00 16.34	
ATOM	2201		GLY	295	3.122	-8.117	3.200 2.822	1.00 19.13 1.00 20.78	
ATOM	2202	0 N	GLY THR	295 296	3.769 3.607	-7.137 -9.354	3.189	1.00 20.78	
ATOM	2204	CA	THR	296	4.909	-9.646	2.618	1.00 10.24	
ATOM	2205	c	THER	296		-10.381	1.319	1.00 15.86	
ATOM	2206	. 0	THR	296		-11.346	1.293	1.00 17.65	
ATOM	2207	СВ	THR	296		-10.506	3.557	1.00 13.92	
ATOM	2208	0G1		296	5.654		4.899	1.00 20.85	
ATOM	2209	CG2		296		-10.411	3.153	1.00 6.06	
MOTA	2210	N	TYR	297	5.213	-9.884	0.235	1.00 11.45	
MOTA	2211	CA	TYR	297	5.005		-1.089	1.00 10.22	
ATOM.	2212	С	TYR	297	6.292		-1.897	1.00 12.07	
MOTA	2213	0	TYR	297	7.174	-9.691	-1.737	1.00 10.00	
MOTA	2214	CB	TYR	297	4.014	-9.562	-1.871	1.00 10.72	
MOTA	2215	·CG	TYR	297	2.669	-9.442	-1.207	1.00 10.66	
MOTA	2216	CD1	TYR	297	1.758	-10.487	-1.259	1.00 11.37	
MOTA	2217	CD2	TYR	297	2.332	-8.315	-0.460	1.00 12.31	
ATOM	2218	CE1	TYR	297	0.546	-10.416	-0.576	1.00 12.44	
MOTA	2219	CE2	TYR	297	1.125	-8.243	0.225	1.00 13.17	
MOTA	2220	CZ	TYR	297	0.236	-9.301	0.165	1.00 19.37	
ATOM	2221	OH	TYR	297	-0.956	-9.274	0.854	1.00 21.32	
ATOM	2222	N	ASP	298		-11.523	-2.780	1.00 8.38	
MOTA	2223	CA	ASP	298		-11.777	-3.721	1.00 7.18	
ATOM	2224	C	ASP	298		-11.149	-5.042	1.00 15.56	
ATOM	2225	0	ASP	298		-11.728	-5.777	1.00 17.52	
ATOM	2226	CB	ASP	298		-13.286	-3.923	1.00 8.20	
ATOM	2227	CC	ASP ·	298		-13.647	-4.958	1.00 13.40	
atom Atom	2228 2229	OD1		298		-12.771	-5.468	1.00 12.72	
ATOM	2229	OD2		298		-14.855	-5.259	1.00 19.52 1.00 10.89	
ATOM ATOM	2230	N	LEU	299	7.546	-9.975 -9.228	-5.334		
ATOM ATOM	2231	CA C	LEU	299 299	7.209 7.509	-9.228 -9.976	-6.539 -7.813	1.00 7.89 1.00 8.02	
ATOM	2232	0	LEU	299	6.818	-9.799	-8.807	1.00 8.02	
ATOM	2234		LEU	299 299	7.909	-9.799 -7.871	-6.553	1.00 7.31	
ATOM	2235	cc	LEU	299	7.540	-6.923	-5.408	1.00 10.69	
	223	-0		633	7.340	-0.923	-3.400	7.00 10.03	

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ATOM
          2236
                CD1 LEU
                          299
                                     8.295 -5.624
                                                     -5.553
                                                             1.00 9.61
  HOTA
          2237
                CD2 LEU
                          299
                                     6.031 -6.681
                                                     -5.396
                                                            1.00 12:99
  MOTA
          2238
                N
                    LYS
                          300
                                     8.534 -10.816
                                                     -7.778
                                                            1.00
  ATOM
          2239
                CA
                    LYS
                          300
                                     8.903 -11.599
                                                     -8.948
                                                            1.00 6.84
  MOTA
          2240
                C
                    LYS
                          300
                                     7.714 -12.453 -9.355
                                                            1.00 14.89
  MOTA
          2241
                0
                    LYS
                          300
                                     7.308 -12.472 -10.518
                                                            1.00 15.15
  ATOM
          2242
                CB
                    LYS
                          300
                                    10.091. -12.495
                                                    -8.624
                                                            1.00 9.49
  ATOM
          2243
                CG
                    LYS
                          300
                                    10.488 -13.488
                                                    -9.710
                                                            1.00 27.80
  ATOH
          2244
                CD
                    LYS
                          300
                                    11.752 -14.226
                                                    -9.292
                                                            1.00 44.37
  ATOM
          2245
                CE
                    LYS
                          300
                                    12.067 -15.433 -10.159
                                                             1.00 56.70
  ATOH
          2246
                NZ
                    LYS
                          300
                                    13.235 -16.166
                                                            1.00 65.57
                                                    -9.598
  MOTA
          2247
                N
                          301
                    SER
                                     7.126 -13.101
                                                     -8.359
                                                            1.00 12.97
  ATOM
          2248
                CA
                    SER
                          301
                                     5.968 -13.955
                                                    -8.553
                                                            1.00 13.55
  MOTA
          2249
                C
                    SER
                          301
                                                     -8.866
                                     4.730 -13.125
                                                            1.00 20.42
  MOTA
          2250
                0
                    SER
                          301
                                     4.186 -13.198
                                                    -9.976
                                                            1.00 21.11
 MOTA
          2251
               CB
                    SER
                          301
                                     5.715 -14.775
                                                    -7.290
                                                            1.00 16.92
 ATOM
         2252
               œ
                   SER
                          301
                                     4.473 -15.458
                                                    -7.351 1.00 27.19
 ATOM
         2253
               N
                   VAL
                          302
                                    4.330 -12.294
                                                    -7.901
                                                            1.00 16.40
 MOTA
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MOTA	2428	N	LEU	327	22.724		-16.594	1.00 14.79
	2429	CA	LEU	327	22.111	•	-15.317	1.00 14.65
MOTA	2430	C	LEU	327	20.612		-15.383	1.00 23.21
MOTA	2431	0	LEU	327	20.002		-16.439	1.00 25.97
ATOM	2432	CB	LEU	327	22.346		-14.992	1.00 13.29
ATOM	2433	CG	LEU	327	21.934		-13.604	1.00 15.65
ATOM	2434	CD1		327	22.728	-7.487		1.00 16.46
ATOM	2435	CD2		327	22.180	-5.219 -9.268		1.00 14.83
MOTA MOTA	2436 2437	N. CA	LYS	328 328	20.022 18.595	-9.530		1.00 13.80
ATOM	2438	CA	LYS LYS	328	18.190		-12.714	1.00 12.00
ATOM	2439	0	LYS	328		-9.401		1.00 14.81
ATOM	2440	CB	LYS	328		-10.945		1.00 12.41
ATOM	2441	CG	LYS	328		-12.020		1.00 15.34
ATOM	2442	CD	LYS	328		-13.387		1.00 23.38
ATOM	2443	CE	LYS	328		-14.531		1.00 34.78
ATOM	2444	NZ	LYS	328		-15.847		1.00 52.92
ATOM	2445	N	LEU	329	16.893	-9.193		1.00 14.22
ATOM	2446	CA	LEU	329	16.362	-8.995		1.00 13.57
MOTA	2447	С	LEU	329		-10.323 -		1.00 18.65
MOTA	2448	0	LEU	329		-10.884	-10.685	1.00 17.39
MOTA	2449	CB	LEU	329	15.143	-8.079	-11.184	1.00 12.21
MOTA	2450	CG	LEU	329	14.728	-7.426	-9.873	1.00 14.57
MOTA	2451	CD1	LEU	329	15.929	-6.837	-9.172	1.00 13.69

		_					-10.180	1.00 14.70
ATOM	2452		LEU	329	13.705		-9.680	1.00 15.18
ATOM	2453	N	SER	330		-10.809 -12.071		1.00 14.75
ATOM	2454	CX	SER	330	16.847		-7.766	1.00 19.86
MOTA	2455	С	SER	330		-11.963	-7.482	1.00 22.85
MOTA	2456	0	SER	330		-12.879		1.00 21.05
MOTA	2457		SER	330		-12.532	-8.525	
MOTA	2458	OG	SER	330		-12.336	-9.544	1.00 39.22
MOTA	2459	N	LYS	331	16.011	-10.838	-7.067	1.00 13.26
MOTA	2460	CX	LYS	331		-10.653	-5.880	1.00 11.98
MOTA	2461	С	LYS	331	14.383	-9.353	-5.953	1.00 12.78
ATOM	2462	0	LYS	331		-8.310		-1.00 11.98
ATOM	2463	CB	LYS	331		-10.668	-4.622	1.00 13.85
MOTA	2464	CC	LYS	331		-11.219	-3.355	1.00 13.58
ATOM	2465	CD	LYS	331		-12.736	-3.420	1.00 14.30
MOTA	2466	CE	LYS	331		-13.191		1.00 17.75
MOTA	2467	NZ	LYS	331		-12.980	-1.115	1.00 46.06
MOTA	2468	N	ALA	332	13.095	-9.444	-5.651	1.00 8.06
MOTA	2469	CA	ALA	332	12.222	-8.274		
MOTA	2470	С	ALA	332	11.167		-4.572	1.00 11.68
MOTA	2471	0	ALA	332	10.302	-9.471	-4.785	1.00 9.24
ATOM	2472	CB	ALA	332	11.591		-6.998	1.00 8.18
MOTA	2473	N	VAL	333	11.309	-8.026	-3.394	1.00 9.58
MOTA	2474	CA	VAL	333	10.394	-8.297	-2.294	1.00 9.85
ATOM	2475	С	VAL	333	9.725	-7.032	-1.758	1.00 13.40
ATOM	2476	0	VAL	333	10.352	-5.973	-1.684	1.00 12.46
ATOM	2477	CB	VAL	333	11.137	-9.022	-1.132	1.00 13.78
MOTA	2478	CG1	VAL	333	10.172	-9.353	0.014	1.00 12.99
MOTA	2479	CG2	VAL	333		-10.290	-1.645	1.00 13.83
ATOM	2480	N	HIS	334	8.441	-7.152	-1.427	1.00 9.41
MOTA	2481	CA	HIS	334	7.669		-0.873	1.00 9.17
MOTA	2482	C	HIS .	334		-6.514	0.469	1.00 15.24
MOTA	2483	0	HIS	334	6.784	-7.688	0.632	1.00 15.50
ATOM	2484	CB	HIS	334	6.493	-5.694	-1.787	1.00 9.35
MOTA	2485	CG	HIS	334	5.638	-4.579	-1.254	1.00 11.86
ATOM	2486	ND1	HIS	334	5.966	-3.245	-1.391	1.00 13.40
ATOM	2487	CD2	HIS	334	4.461	-4.609	-0.593	1.00 12.02
ATOM	2488	CE1	HIS	334	5.025	-2.501	-0.834	1.00 12.10
ATOM	2489	NE2	HIS	33,4	4.101	-3.309	-0.343	1.00 11.99
MOTA	2490	N	LYS	335 .	7.064	-5.592	1.427	1.00 12.37
MOTA	2491	CA	LYS	335	6.505	-5.848	2.752	1.00 11.91
ATOM	2492	C	LYS	335	5.866	-4.565	3.239	1.00 15.10
MOTA	2493	0	LYS	335	6.538	-3.542	3.391	1.00 15.49
MOTA	2494	CB	LYS	335	7.562	-6.311	3.756	1.00 15.51
MOTA	2495	CC	LYS	335	6.958	-6.733	5.106	1.00 37.80
MOTA	2496	CD -	LYS	335	8.018	-7.241	6.074	1.00 44.20
MOTA	2497	CE	LYS	335	7.448	-7.512		1.00 53.49
MOTA	2498	NZ	LYS	335	6.509	-8.655	7.395	1.00 71.81
MOTA	2499	N	ALA	336	4.550	-4.598	3.387	1.00 11.02
MOTA	2500	CA	ALA	336	3.801	-3.441	3.845	1.00 10.26
MOTA	2501	С	ALA	336	3.301	-3.726	5.242	1.00 13.86
MOTA	2502	O .	ALA	336	2.871	-4.846	5.520	1.00 12.29
MOTA	2503	CB	ALA	336	2.619	-3.179	2.912	1.00 10.63
MOTA	2504	N	VAL	337	3.422	-2.739	6.126	1.00 12.67
MOTA	2505	CA	VAL	337	2.950	-2.861	7.502	1.00 14.10

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ATO			C VA			2.0		-1. 6 6	4 7.	756	1.0	0 1	19.	06
ATO			O VA			2.3		-0.52		411	1.0	0 1	9.	09
ATO	•		B VA			4.1		-2.80	6 8.	540				
ATON			G1 VA			3.6		-3.22	0 9.	913				
ATON			G2 VAI			5.2		-3.69		119	1.0	0 1	9.1	80
ATOH			LE			0.8		-1.93		285	1.0			
ATOM			A LE			-0.1		-0.88	9 8.6	503	1.0	0 1	0.6	58
ATOM						-0.64		-1.13			1.0	0 1	5.5	52
ATOM						-1.22		-2.18			1.0	0 1	4.2	26
ATOM			B LEU			-1.30		-0.90		524	1.00)	9.0)1
ATOM ATOM			G LEU			-2.50		-0.011			1.00	1	0.3	3
			D1 LEU			-2.40		1.321			1.00		8.7	
MOTA MOTA			D2 LEU			-3.78		-0.727			1.00			
ATOM						-0.45		-0.142			1.00			
ATOM	2520				•	-0.91		-0.200			1.00			
ATOM	2523 2523					-2.02		0.826			1.00			
ATOH	2523					-1.81		2.023			1.00			
ATOM	2524			339		0.23		0.086			1.00			
ATOM	2525		31 THR 32 THR	339		1.21		-0.959			1.00			
ATOM	2526			339		-0.28		0.161			1.00			
ATOM	2527		ILE ILE	340		-3.20		0.331	12.8		1.00			
ATOM	2528		ILE	340 340		-4.389		1.141	13.13		1.00			
ATOM	-2529	-	ILE			-4.735		1.064	14.6		1.00			
ATOM	2530			340 340 -		-4.795		-0.032	15.17		1.00			
ATOM	2531			340		-5.633 -5.412		0.620	12.34		1.00			
ATOM	2532	CG		340		-6.903		0.751	10.83		1.00			
ATOM	2533			340		-5.170		1.377	12.75		1.00			
ATOM	2534	N	ASP	341		-4.932		2.154	10.39		1.00			
ATOM	2535	CA		341		-5.310		2.292	15.25 16.67		1.00			
ATOM	2536	C	ASP	341		-6.199		3.503	16.94		1.00			
MOTA	2537	0	ASP	341		-6.821		4.008	16.00		1.00			
MOTA	2538	CB	ASP.	341		-4.097		2.221	17.63		1.00			
ATOH :	2539	CG	ASP	341		-3.190		3.437	17.54		1.00			
ATOM	2540	OD:	LASP	341		-3.500		4.422	16.85		1.00			
ATOM	2541	OD:	2 ASP	341		-2.135		3.389	18.21		1.00			
MOTA	2542	N	GLU	342		-6.250		3.963	18.19		1.00			
ATOM	2543	CA	GLU	342		-7.093		5.095	18.59		.00			
ATOM	2544	С	GLU	342		-6.603		5.439	18.049		.00			
MOTA	2545	0	GLU	342		-7.419		7.318	17.762		.00			
ATOM	2546	CB	GLU	342		-7.283		5.157	20.111		.00			
MOTA	2547	CG	GLU	342		-8.017	3	.973	20.724		.00 2			
MOTA	2548	CD	GLU	342		-7.121	2	.773	21.007		.00 3			
MOTA	2549		GLU	342		-5.878	2	.880	20.936		.00 3			
MOTA	2550		GLU	342		-7.674	1	.705	21.318		.00	1.0		
NOOM	2551	N	LYS	343		-5.282	6	. 595	17.934		.00 2			
MOTA	2552	CA	LYS	343		-4.637	7	.783	17.359		.00 2			
MOTA	2553	С	LYS	343		-3.143	7	.809	17.574		.00 3			
MOTA	2554	0	LYS	343		-2.689			18.638		.00 3			
MOTA	2555	CB	LYS	343		-5.202			17.873		.00 2			
MOT	2556	CG	LYS	343		-4.622			17.113	1	.00 4	3.2	2	
TOM	2557	CD	LYS	343		-5.145			17.618	1.	.00 5	9.1	4	
TOM	2558	CE	LYS	343		-4.536			16.854	1.	.00 8	1.8	7	
MOT	2559	NZ	LYS	343		-5.391	14.	.019	16.935		00 9			

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ATOM	2560		GLY	344	-2.393	7.401	16.548	1.00 36.10
ATOM	2561	. CA	GLY	344	-0.940	7.390	16.605	1.00 37.91
ATOM	2562	C	GLY	344	-0.393	7.219	18.003	1.00 47.44
MOTA	2563		GLY	344	-0.635	6.198	18.656	1.00 50.02
MOTA	2564	N	THR	345	0.215	8.279	18.517	1.00 44.73
ATOM	2565	CA	THR	345	0.782	8.240	19.853	1.00 45.38
MOTA	2566	C	THR	345	0.267	,9.472	20.581	1.00 48.09
MOTA	2567	0	THR	345	-0.049	10.480	19.946	1.00 47.67
MOTA	2568	CB	THR	345	2.313	8.273	19.797	1.00 62.95
ATOM	2569	· OG	1 THR	345	2.741	9.424	19.051	1.00 67.56
MOTA	2570	CG	2 THR	345	2.840	7.014	19.122	1.00 62.12
MOTA	2571	N	GLU	346	0,163	9.387	21.904	1.00 44.03
MOTA	2572	CA	GLU	346	-0.325	10.518	22.679	1.00 43.30
ATOM	2573	С	GLU	346	0.562	11.738	22.461	1.00 46.46
ATOM	2574	0	GLU	346	1.746	11.732	22.814	1.00 47.39
ATOM	2575	СВ	GLU	346	-0.401		24.165	1.00 45.00
ATOM	2576	CG	GLU	346	-1.182	11.212	24.978	1.00 64.69
ATOM	2577	CD	GLU	346	-1.166	10.927	26.475	1.00108.58
ATOM	2578	OE:	1 GLU	346	-0.692	9.845	26.897	1.00110.00
ATOM	2579	OE:		346	-1.632	11.799	27.237	1.00110.00
ATOM.	2580	N	ALA	347	-0.026	12.754	21.832	1.00 40.86
ATOM	2581	CA	ALA	347	0.634	14.021	21.536	1.00 39.63
ATOM	2582	С	ALA	347	-0.460	15.092	21.567	1.00 41.81
ATOM	2583	0	ALA	347	-1.499	14.885	22.205	1.00 42.33
ATOM	2584	СВ	ALA	347	1.296	13.970	20.159	1.00 40.47
MOTA	2585	N	ALA	348	-0.235	16.231	20.879	1.00 35.47
MOTA	2586	CA	ALA	348	-1.254	17.271	20.847	1.00 34.53
ATOM	2587	С	λLA	348	-2.568	16.718	20.362	1.00 41.09
MOTA	2588	0	ALA	348	-2.750	16.513	19.173	1.00 41.40
ATOM	2589	CB	ALA	348	-0.778	18.419	19.934	1.00 34.78
MOTA	2590	N	GLY	349	-3.484	16.471	21.320	1.00 38.15
ATOM	2591	CA	GLY	349	-4.744	15.826	20.982	1.00 38.10
MOTA	2592	С	GLY	349	-5.574	16.643	20.033	1.00 41.35
MOTA	2593	0	GLY	349	-6.114	16.085	19.091	1.00 42.51
MOTA	2594	N	ALA	350	-5.673	17.965	20.291	1.00 34.71
MOTA	2595	CA	ALA	350	-6.489	18.821	19.442	1.00 33.31
MOTA	2596	C	ALA	350	-7.930	18.421	19.594	1.00 36.83
MOTA	2597	0	ALA,	350	-8.424	17.591	18.846	1.00 37.89
MOTA	2598	CB	ALA	350	-6.018	18.750	17.975	1.00 33.89
MOTA	2599	N	MET	351	-8.607	19.017	20.593	1.00 30.79
MOTA	2600	CA	MET	351	-9.956	18.566	20.888	1.00 29.73
MOTA	2601	С	MET	351	-10.993	19.259	20.050	1.00 31.90
MOTA	2602	0	MET	351	-11.024	20.477	19.983	1.00 32.74
MOTA	2603	CB	MET	351	-10.280	18.653	22.394	1.00 32.16
MOTA	2604	CG	MET	351	-9.025	18.414	23.262	1.00 36.44
MOTA	2605	SD	MET	351	-8.109	16.934	22.720	1.00 41.73
ATOM	2606	CE	MET	351	-9.418	15.696	22.959	1.00 37.83
ATOM	2607	N	PHE	352	-11.848	18.447	19.401	1.00 25.13
ATOM	2608	CA	PHE	352	-12.892	19.029	18.577	1.00 23.52
MOTA	2609	С	PHE	352	-14.245	18.711	19.149	1.00 28.36
MOTA	2610	0	PHE	352	-14.342		20.150	1.00 30.86
MOTA	2611	CB	PHE	352	-12.778		17.131	1.00 25.09
MOTA	2612	CG	PHE	352	-12.827		17.095	1.00 26.88
ATOM .	2613	CD1		352	-11.641		17.183	1.00 29.01
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ATOM	2722		LYS	365	-20.930	-4.371	16.320		_
MOTA	2723	CE	LYS	365	-22.01	-4.001	17.311		
ATOM	2724		LYS	365	-22.746	-5.207	17.780)
ATOM	2725	N,	PHE	366	-17.081	-2.979	12.164	1.00 4.94	1
ATOM	2726	CA	PHE	366	-16.163	-3.613	11.226	1.00 5.21	1
ATOM	2727	С	PHE	366	-15.707		11.942	1.00.11.04	1
MOTA	2728	0	PHE	366	-14.558	-4.973	12.395	1.00 9.49	j
MOTA	2729	CB	PHE	366	-14.990	-2.678	10.860		5
MOTA	2730	CG	PHE	366	-15.339	-1.691	9.761)
ATOM	2731	CD	1 PHE	366	-16.106	-0.565	10.039	1.00 10.00)
MOTA	2732	CD:	2 PHE	366	-15.010	-1.954	8.436	1.00 8.12	!
MOTA	2733	CE.	1 PHE	366	-16.542	0.259	9.020	1.00 10.80)
MOTA	2734	CE:	2 PHE	366	-15.448	-1.126	7.414	1.00 9.85	,
ATOM	2735	CZ	PHE	366	-16.213		7.706	1.00 8.09	į
MOTA	2736	N	ASN	367	-16.675	-5.781	12.090	1.00 10.96	,
ATOM	2737	CA	ASN	367	-16.511	-7.057	12.775	1.00 12.66	
ATOM	2738	С	ASN	367	-16.575	-8.273	11.863	1.00 20.63	
MOTA	2739	.0	ASN	367	-17.025	-9.343	12.274	1.00 23.39	
ATOM	2740	CB	ASN	367	-17.570	-7.211	13.873	1.00 14.33	
MOTA	2741	CG	ASN	367	-18.982	-7.299	13.321	1.00 53.89	
ATOM	2742		ASN	367	-19.227	-7.000	12.150	1.00 45.61	
MOTA	2743	ND	2 ASN	367	-19.923	-7.702	14.166	1.00 57.43	
MOTA		N	LYS	368	-16.183	-8.100	10.612	1.00 15.70	
ATOM	2745	CA	LYS	368	-16.163	-9.201	9.648	1.00 15.30	
MOTA	2746	C	LYS	368	-15.231	-8.792	8.508	1.00 23.90	
ATOM	2747	0	LYS	368	-14.900	-7.595	8.372	1.00 25.23	
MOTA	2748	CB	LYS	368	-17.577	-9.530	9.149	1.00 14.67	
ATOM	2749	CG	LYS	368	-18.279	-8.405	8.423	1.00 5.03	
MOTA	2750	CD	LYS	368	-19.624	-8.855	7.894	1.00 2.48	
ATOM	2751	CE	LYS	368	-20.409	-7.697	7.293	1.00 26.14	
ATOM	2752	NZ	LYS	368	-21.767	-8.081	6.805	1.00 49.97	
ATOM	2753	N	PRO	369	-14.826	-9.751	7.652	1.00 18.79	
ATOM	2754	CA	PRO	369	-13.927	-9.382	6.567	1.00 17.06	
ATOM	2755	C	PRO	369	-14.315	-8.087	5.846	1.00 15.58	
ATOM	2756	0	PRO	369	-15.498	-7.832	5.590	1.00 13.98	
MOTA	2757	СВ	PRO	369			5.682	1.00 18.58	
MOTA	2758	CG	PRO	369	-14.067	-11.718	6.699	1.00 22.28	
ATOM	2759	CD	PRO	369	-15.202		7.525	1.00 17.81	
MOTA	2760	N	PHE	370	-13.335	-7.198	5.698	1.00 10.74	
ATOM	2761	CA	PHE	370	-13.554	-5.924	4.999	1.00 9.37	
ATOM	2762 2763	C	PHE	370	-12.338	-5.508	4.178	1.00 7.98	
ATOM		0	PHE	370	-11.206	-5.867	4.520	1.00 6.62	,
ATOM ATOM	2764 2765	CB CG	PHE	370	-13.946	-4.795	5.979	1.00 10.78	
ATOM	2766		PHE	370	-12.879 -11.902	-4.437	6.990	1.00 10.60	
ATOM	2767		PHE	370 370	-12.861	-3.492 -5.047	6.695	1.00 12.76	
	2768		PHE	370			8.236	1.00 11.98	
ATOM	2769		PHE		-10.918 -11.888	-3.157	7.629	1.00 13.14	
ATOM	2770	CZ	PHE	370 370	-10.910	-4.725 -3.779	9.176	1.00 15.00	
ATOM	2771	N	VAL	370 371	-12.564		8.872	1.00 13.05	
ATOM	2772	CA	VAL	371	-11.444	-4.811 -4.329	3.067	1.00 3.46	
ATOM	2773	CA	VAL	371 371	-11.239	-4.329	2.256	1.00 4.17	
ATOM	2774		VAL		-12.083		2.601	1.00 6.84	
ATOM	2775		VAL	371 371	-11.695	-2.249	3.258	1.00 6.12	
WI OU	2113	CB	AVT	371	-11.033	-4.504	0.720	1.00 9.05	

ATOM	277	6 0	G1 VA	L 371		-12.22	21 -5.90	0 0.4	25 1.00	9.32
MOTA	277		G2 VA	L 371		-12, 62				
ATOM	277	8 N				-10.13	2 -2.25			-
ATOM	277	9 C	A PH	Ė 372		-9.88			-	8.81
ATOM	278	_	PHI	E 372		-8.70	8 -0.28			10.01
ATOM	278					7.89				10.68
MOTA	278		B PHI	E 372		-9.61				12.58
MOTA	278		G PH	372		-8.35				15.84
MOTA	278		D1 PHE			-8.30				21.46
ATOM	278		D2 PHE			-7.18	9 -0.54			18.60
ATOM	2786		El PHE			-7.12	2 -3.28			22.91
ATOM	2787		E2 PHE		•	-5.99	7 -1.15			22.97
ATOM	2788		_			-5.96				22.13
ATOM	2789					-8.61	2 1.04			4.88
ATOM	2790					-7.50	9 1.68	7 0.89	5 1.00	4.91
MOTA	2791		LEU			-7.12	5 2.94			9.95
ATOM	2792		LEU			-7.918				9.65
ATOM	2793					-7.912		-0.53		4.96
ATOM	2794			•		-8.335	1.061	-1.57	9 1.00	
ATOM	2795		1 LEU			-9.789		-1.39		
ATOM	2796		2 LEU			-8.135	1.658	-2.94		
ATOM	2797		MET	374		-5.882		1.46		9.56
ATOM	2798			374		-5.394			3 1.00 1	
ATOM	2799		MET			-4.968		0.890		
ATOM	2800	. 0	MET	374		-4.134		0.083	2, 1.00 1	2.10
ATOM	2801	CB		374		-4.215			1.00 1	5.63
ATOM	2802	CG		374		-4.611			1.00 2	0.55
ATOM ATOM	2803	SD		374		-5.417				
ATOM	2804	CE	MET	374		-6.308	2.489			
ATOM	2805 2806	N,	ILE	375		-5.608	6.600			
ATOM	2807	CA C	ILE	375		-5.376	7.518			
ATOM	2808		ILE	. 375		-4.724		0.112		
ATOM	2809	CB	ILE ILE	375		-5.187	9.404	1.076		
ATOM	2810	CG		375		-6.727		-1.039		
ATOM	2811	CG2		375 375		-7.222	6.742	-1.928		
ATOM	2812		ILE	375		-6.589	9.148	-1.867		
ATOM	2813	N	GLU	375		-7.874 -3.656	5.633	-1.180		
ATOM	2814	CA	GLU.	376		-3.002	9.238	-0.563	1.00 14	
ATOM	2815	c	GLU	376		-3.920	10.503	-0.212	1.00 14	
ATOM	2816	ō	GLU	376		-4.048	11.596	-0.767	1.00 19	
ATOM	2817	СВ	GLU	376		-1.603	11.755			
MOTA	2818	CG	GLU	376		-0.710	10.607 11.742	-0.842	1.00 16	
MOTA	2819	CD	GLU	376		-1.097	13.127	-0.291	1.00 21	
MOTA	2820		GLU	376		-1.086		-0.798	1.00 33	
MOTA	2821		GLU	376		-1.377	13.331	-2.033	1.00 18	
MOTA	2822	N	GLN	377		-4.550	14.023 12.337	0.021	1.00 13	
MOTA	2823	CA	GLN	377		-5.492	13.396	0.136	1.00 19	
MOTA	2824	C	GLN	377		-5.247	14.235	-0.202	1.00 20	
MOTA	2825	ō	GLN	377		-6.191	14.510	-1.451 -2.197	1.00 27	
MOTA	2826	СВ	GLN	377		-5.676	14.337	0.985	1.00 27	
MOT	2827	CG	GLN	377		-6.428	13.737	2.162	1.00 22 1.00 53	
MOT	2828	CD	GLN	377		-6.529	14.707	3.319	1.00 88	
MOTA	2829	OE1		377		-6.140	15.874	3.205	1.00 88	
								J. EUJ	1.00 09	

	•								
ATOM	283	0 N	E2 GLN	377		-7.04	4 14.239	5 4.44	2 1.00 87.27
ATOM	283	1 N	ASN	378		-3.99			
ATOM	283	2 C		••		-3.65			
ATOM	283	-	ASN	378		-3.53			
ATOM	283	6 0	ASN	378		-4.11			
ATOM	2839	C C		378		-2.37			
ATOM	2836	s co		378		-2.52			
ATOM	2837	O	1 ASN	378		-3.25			
ATOM	2838		2 ASN	378		-1.84			
ATOM	2839	N	THR	379		-2.750			
ATOM	2840	CA	THR	379		-2.496		_	
ATOH.	2841	. с	THR	379		-3.531			
ATOM	2842	0	THR	379		-3.682			
ATOM	2843	CB	THR	379		-1.141			
ATOM	2844	OG	1 THR	379		-1.168	11.474		
ATOM	2845	CG	2 THR	379		-0.078			
MOTA	2846	N	LYS	380		-4.190			
ATOM	2847	CA	LYS	380		-5.186			1.00 11.20
MOTA	2848	С	LYS	380		-4.569			1.00 14.14
ATOM	2849	0	LYS	380		-5.264		-5.507	•
ATOM	2850	CB	LYS	380		-6.359			1.00 13.33
ATOM	2851	CG	LYS	380		-7.021		-5.076	1.00 43.92
MOTA	2852	CD	LYS	380		-8.294		-5.834	1.00 62.33
MOTA	2853	CE	LYS	380		-8.943		-5.320	1.00 75.41
MOTA	2854	NZ	LYS	380		-10.179	14.084	-6.069	1.00 81.22
MOTA	2855	N	SER	381		-3.267	9.033	-4.806	1.00 10.76
ATOM	2856	CA	SER	381		-2.501	7.829	-5.078	1.00 10.64
ATOM	2857	С	SER	381		-2.826	6.773	-4.043	1.00 17.98
MOTA	2858	0	SER.	381		-2.763	7.035	-2.825	1.00 18.86
ATOM	2859	CB	SER	381		-1.009	8.137	-5.013	1.00 12.65
MOTA	2860	OG	SER	381		-0.681	9.193	-5.890	1.00 24.22
ATOM	2861	N	PRO	382		-3.165	5.556	-4.498	1.00 15.96
MOTA	2862	CA	PRO	382		-3.484	4.522	-3.516	1.00 15.23
MOTA	2863	C	PRO	382	•	-2.229	4.026	-2.787	1.00 15.86
atom Atom	2864	0	PRO	382		-1.428	3.275	-3.343	1.00 13.97
ATOM	2865	CB	PRO	382	•	-4.191	3.461	-4.359	1.00 16.74
ATOM	2866 2867	CG.	PRO	382		-3.507	3.573	-5.698	1.00 21.91
ATOM	2868	CD	PRO	382		-3.367	5.069	-5.880	1.00 17.13
ATOM	2869	N CA	LEU	383		-2.046	4.542	-1.572	1.00 11.15
ATOM	2870	CA	LEU LEU	383 383		-0.918 -0.941	4.229	-0.704	1.00 10.25
ATOM	2871	0	LEU	383		0.078	2.803 2.122	-0.201 -0.202	1.00 15.69
ATOM	2872	СВ	LEU	383		-0.886	5.190	0.494	1.00 17.23
MOTA	2873	CG	LEU	383		-0.330	6.617	0.332	1.00 9.55 1.00 12.09
MOTA	2874		LEU	383		-0.693	7.455	1.530	1.00 12.09
MOTA	2875		LEU	383		1.174	6.596	0.129	1.00 12.37
MOTA	2876	N	PHE	384		-2.105	2.345	0.221	1.00 9.64
MOTA	2877	CA	PHE	384		-2.237	0.991	0.733	1.00 7.84
MOTA	2878	С	PHE	384		-3.581	0.451	0.283	1.00 13.29
MOTA	2879	0	PHE	384		-4.559	1.202	0.192	1.00 14.82
MOTA	2880	СВ	PHE	384		-2.279	0.989	2.272	1.00 8.59
MOTA	2881	CG	PHE	384		-0.933	1.120		1.00 9.77
MOTA	2882	CD1		384		-0.204	-0.013		1.00 12.34
MOTA	2883	CD2	PHE	384		-0.418	2.367		1.00 11.86

ATOR	1 288	34 (E1 PH	E 384	1.004 0.085 3.979 1.00 12.93
ATOM	288		E2 PH		1.00 12.3
ATOM	1 . 288		Ż PHI		1.00 14.00
ATOM	288	7 N	MET		1.00
ATOM	288	8 C	A MET	385	
ATOM	288	9 0	MET		-4.769 -2.816 0.368 1.00 5.90
ATOM	289	0 0	. MET		-3.680 -3.368 0.518 1.00 2.53
ATOM	289	1 0	B MET		-5.069 -1.722 -1.877 1.00 8.34
ATOM		2 C	G MET		-6.406 -2.403 -2.150 1.00 11.63
ATOM		3 S	D MET	385	-7.016 -2.177 -3.806 1.00 15.95
ATOM			E MET	385	-8.683 -2.637 -3.625 1.00 12.69
MOTA			GLY	386	-5.890 -3.290 0.887 1.00 2.67
ATOM			A GLY	386	-5.870 -4.544 1.597 1.00 2.61
ATOM	2891		GLY	386	-7.235 -5.010 2.027 1.00 8.66
ATOM	2898		GLY	386	-8.228 -4.350 1.761 1.00 8.72
ATOM	2899		LYS	387	-7.274 -6.176 2.658 1.00 9.06
ATOM	2900			387	-8.501 -6.778 3.168 1.00 9.44
MOTA		_	LYS	387	-8.129 -7.460 4.468 1.00 14.80
MOTA	2902			387	-7.091 -8.115 4.559 1.00 14.15
ATOM	2903			387	-9.067 -7.823 2.195 1.00 10.21
ATOM	2904			387	-10.168 -8.686 2.800 1.00 10.53
ATOM	2905			387	-10.771 -9.628 1.789 1.00 17.82
ATOM	2906			387	-9.741 -10.614 1.280 1.00 30.69
ATOM	2907	_		387	-10.341 -11.615 0.359 1.00 46.30
ATOM ATOM	2908		VAL	388	-8.937 -7.231 5.493 1.00 12.28
ATOM	2909	CA		388	-8.718 -7.849 6.782 1.00 11.26
ATOM	2910	C	VAL	388	-9.748 -8.959 6.904 1.00 14.50
ATOM	2911 2912	0	VAL	388	-10.952 -8.720 6.825 1.00 12.32
ATOM	2913	CB	VAL	388	-8.852 -6.834 7.952 1.00 14.78
ATOM	2914		VAL VAL	388	-8.686 -7.539 9.285 1.00 14.30
ATOM	2915	N		388	-7.802 -5.738 7.818 1.00 14.55
ATOM	2916	CA	VAL VAL	389	-9.251 -10.186 6.900 1.00 14.18
ATOM	2917	C	VAL	389	-10.092 -11.356 7.043 1.00 15.05
ATOM	2918	ō	VAL	389 389	-10.113 -11.638 8.545 1.00 23.37
ATOM	2919	CB	VAL	389	-11.181 -11.822 9.135 1.00 25.50
ATOM	2920		VAL	389	-9.535 -12.590 6.237 1.00 18.20
ATOM	2921		VAL	389	-9.645 -12.354 4.747 1.00 17.82 -8.089 -12.863 6.580 1.00 18.27
MOTA	2922	N	ASN	390	
ATOM	2923	CA	ASN	390	
ATOM	2924	C	ASN.	390	8 444
ATOM	2925	0	ASN	390	£ 486 AA AA-
MOTA	2926	СВ	ASN	390	
MOTA	2927	CG	ASN	.390	
MOTA	2928		ASN	390	
NTOM	2929		ASN	390	
NOTA	2930	N	PRO	391	
MOTA	2931	CA	PRO	391	-7.946 -10.034 12.127 1.00 20.72 -6.941 -9.213 12.789 1.00 20.87
MOTA	2932	C	PRO	391	-5.886 -10.027 13.503 1.00 31.97
MOTA	2933	0	PRO	391	-4.750 -9.585 13.641 1.00 33.17
MOTA		CB	PRO	391	-7.776 -8.394 13.765 1.00 21.61
MOT		CG _.	PRO	391	-9.039 -8.187 12.996 1.00 25.72
TOM .		CD	PRO	391	-9.286 -9.624 12.588 1.00 21.28
TOM	2937	N	THR	392	-6.269 -11.230 13.910 1.00 33.77
				•	

```
ATOM
          2938
                CA
                   THR
                          392
                                   -5.388 -12.134 14.630
                                                           1.00 35.56
   ATON
          2939
                Ċ
                                 -4.138 -12.469 13.823
                    THR
                          392
                                                           1.00 .41.19
  ATOM
          2940
               O THR
                          392
                                   -3.042 -12.602 14.379
                                                           1.00 42.51
  ATOM
          2941
              CB THR
                         392.
                                  -6.136 -13.423
                                                   15.015
                                                           1.00 56.77
  ATOM
          2942
               OG1 THR
                        . 392
                                   -7.263 -13.091
                                                   15.843
  ATOM
                                                           1.00.59.19
          2943 CG2 THR
                         392
                                  -5.219 -14.371
                                                   15.764
  ATOM
                                                           1.00 61.71
         2944 N
                   GLN
                         393
                                  -4.299 -12.591
                                                  12.512 1.00 36.59
  ATOM
         2945 CA GLN
                       393 جي
                                  -3.174 -12.891
                                                  11.639
  ATOM
                                                          1.00 36.28
         2946 C
                   GLN
                         393
                                  -2.345 -11.630
                                                  11.429
                                                          1.00 43.21
  MOTA
         2947 0
                   GLN -
                         393
                                  -2.894 -10.531
                                                  11.321
                                                          1.00 44.40
  ATOM
         2948 CB
                  GLN
                         393
                                  -3.684 -13.393
                                                  10.290
                                                          1.00 37.18
  ATOM .
         2949 CG
                  GLN
                         393 .
                                  -2.609 -13.457
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                                                          1.00 38.11
 MOTA
         2950 CD GLN
                         393
                                  -3.165 -13.823
                                                   7.853
                                                         1.00 59.92
 ATOM
         2951
              OE1 GLN
                        393.
                                  -4.385 -13.831
                                                   7.634 1.00 50.05
 MOTA
        2952 NE2 GLN
                        393
                                  -2.275 -14.108
                                                   6.921 1.00 61.15
 ATOM
        2953 UN
                  LYS
                        394
                                -1.029 -11.792
                                                 11.357
                                                         1.00 41.17
 ATOM
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                                                 11.122 1.00 73.90
 ATOM
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              C
                  LYS
                        394
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                                                 10.275 1.00106.52
 ATOM
        2956
              0
                  LYS
                        394
                                  1:137 -12.250
                                                  9.864
                                                        1.00 66.01
 MOTA
        2957~ CB. LYS
                        394
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                                                 12.437
                                                        1.00 76.30
 ATOM
              CG LYS
        2958
                        394
                                  1.231
                                         -8.828
                                                12.249
                                                        1.00 89.36
 ATOM
        2959
              CD LYS
                        394
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                                                13.564
                                                        1.00 97.21
ATOM
        2960
             CE LYS
                        394
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                                                        1.00100.59
        2961 NZ LYS
ATOM
                       394
                                  3.234
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                                                14.594
                                                        1.00101.87
ATOM
        2962
             OXT LYS
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                                 1.924 -10.199
                                                  9.997 - 1.00 66.01
TER
      2963
                · LYS
                       394
HETATM 2964 S
                 SCC-
                         1
                               -22.279
                                          4.447
                                                 -3.715
HETATM 2965 C1 SCC
                                                        1.00 31.26
                         1
                               -23.852
                                                -4.634
                                          4.645
HETATM 2966 C2 SCC
                                                        1.00 27.60
                         1
                               -23.920
                                         5.944
                                                -5.448 1.00 26.84
CONECT 1693 1692 2964
CONECT 2964 1693 2965
CONECT 2965 2964 2966
CONECT 2966 2965
MASTER
            - 0
                            - 0
                               . .0
                                                   2965
END
                                                                    29
```

We claim:

1. A furin endoprotease inhibitor comprising a mimetic compound to that portion of α_1 -antitrypsin Portland that comprises amino acid sequence Arg-Xaa-Xaa-Arg at positions 355 through 358 of the amino acid sequence of α_1 -antitrypsin Portland, wherein the atoms of said mimetic are arranged in three dimensional conformation at positions equivalent to those of the atoms comprising the amino acid sequence Arg-Xaa-Xaa-Arg.

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2. A homogenous composition of matter comprising a mimetic compound to that portion of α_1 -antitrypsin Portland that comprises an amino acid sequence Arg-Xaa-Xaa-Arg at positions 355 through 358 of the amino acid sequence of α_1 -antitrypsin Portland, wherein the atoms of said mimetic are arranged in three dimensional conformation at positions equivalent to those of the atoms comprising the amino acid sequence Arg-Xaa-Xaa-Arg.

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3. A method of blocking endoproteolytic activation of a bacterial toxin comprising the step of contacting a cell in the presence of the toxin with a furin endoprotease inhibitor according to Claim 1.

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4. The method of Claim 3 wherein the bacterial toxin is diphtheria toxin of Corynebacterium diptheriae.

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5. The method of Claim 3 wherein the bacterial toxin is anthrax toxin of Bacillus anthracis.

The method of Claim 3 wherein the bacterial toxin is Pseudomonas

aeruginosa exotoxin.

6.

- 30
- 7. A pharmaceutical composition comprising a therapeutically effective amount of a furin endoprotease inhibitor of Claim 1 and a pharmaceutically acceptable carrier or diluent.

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- 8. A method of inhibiting bacterial infection of cells comprising contacting the cells with a furin endoprotease inhibitor according to Claim 1.
- 9. The method of Claim 8 wherein the bacterial toxin is diphtheria toxin of Corynebacterium diptheriae.
 - 10. The method of Claim 8 wherein the bacterial toxin is anthrax toxin of Bacillus anthracis.
- 10 11. The method of Claim 8 wherein the bacterial toxin is *Pseudomonas* aeruginosa exotoxin.
 - 12. A method of inhibiting viral infection of cells comprising contacting the cells with a furin endoprotease inhibitor according to Claim 1.
 - 13. The method of Claim 12 wherein the virus is cytomegalovirus.
 - 14. A method of blocking endoproteolytic viral protein maturation comprising the step of contacting a cell in the presence of the toxin with a furin endoprotease inhibitor according to Claim 1.
 - 15. The method of Claim 14 wherein the virus is cytomegalovirus.
 - 16. A pharmaceutical composition according to Claim 7, further comprising an antibacterial compound.
 - 17. A pharmaceutical composition according to Claim 7, further comprising an antiviral compound.
- 18. A furin endoprotease inhibitor comprising a mimetic compound having the structure:

C(L1-R1)-E-F-G-H-I-J(L2-R2)

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wherein "C" is a mimetic element that is equivalent to a first alpha carbon;

"J" is a mimetic element that is equivalent to a second alpha carbon; whereby "C" and "J" are conformationally hindered;

"E", "G", and "I" are planar moieties having dimensions substantially similar to a peptide bond;

"F" and "H" are mimetic elements that are each equivalent to a conformationally-hindered alpha carbons;

wherein either "F" or "H" are restricted by being integrally covalently linked to a cyclic planar moiety selected from the group consisting of cyclopentane, cyclopentene, furan, tetrahydrofuran, thiophene, pyrrole, or pyrrolidine, or wherein "F" or "H" is covalently linked to a sterically-hindered group;

and whereby E-F-G-H-I is most preferably substantially planar and deviates from this planar structure by no more than from about 1 to about 20 degrees from said plane and wherein the length of the molecule along the distance between the "C" and "J" components (C-E-F-G-H-I-J) is preferably from about 7.5 to about 11.5 Angstroms;

R1 and R2 are each positively-charged residues;

L1 and L2 are each linker moieties selected from the group consisting of methylenes and mimetic elements equivalent thereto;

wherein R1 and R2 are from about 5 to about 7 Angstroms away from their respective alpha carbon equivalents, "C" and "J";

and whereby R1 and R2 are displaced relative to each other along the longitudinal axis of the molecule to subtend an angle of from about 15 to about 25 degrees.

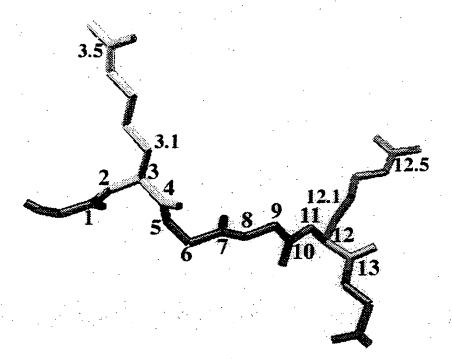


Figure 1A

1 / 9

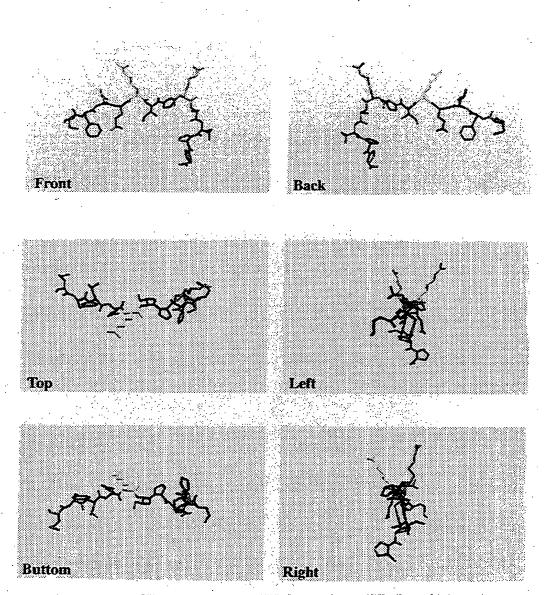
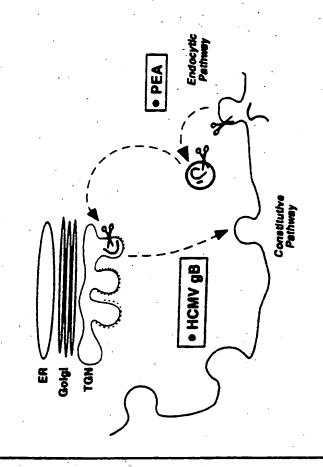


Figure 1B





Pseudomonas axotoxin A

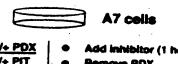
Protoxin from Pseudomonas acuginosa
Cleaved by furth in endosomas
Pseudomonas aeruginosa
Major complication in:
Burn patients
Cyatic fibrosis petients

Pro
HCMV glycoprotein gB

Virton-associated protein
Cleaved by furth

Human Cytomegalovirus
Major cause of morbidity and mortality in immunocompromised individuals transplant and AIDS petients

PEA assay





- · Add PEA (6 M)
- Label with [*S]-Met/Cye (30 min)
- TCA precipitate cellular proteins

Scintillation count

FIGURE 3

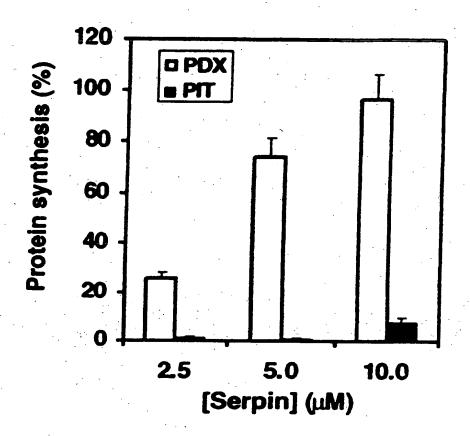


FIGURE 4

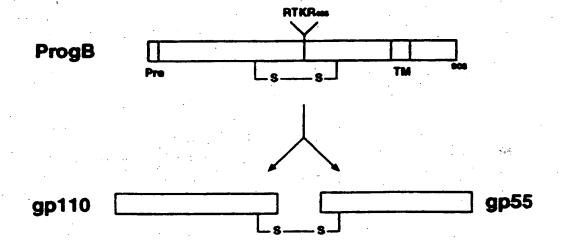


FIGURE 5

HCMV plaque reduction assay

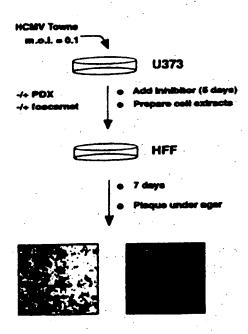


FIGURE 6

Inhibition of HCMV plaque formation by $\alpha_1\text{-PDX}$ and foscernet

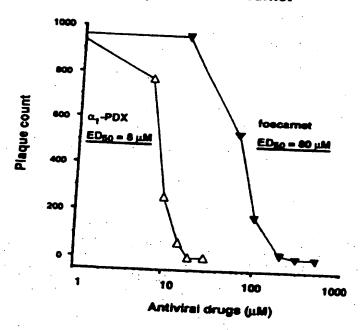


FIGURE 7

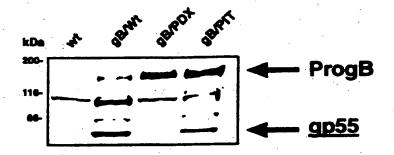


FIGURE 8

SEQUENCE LISTING

(1) GENERAL INFORMATION:

- (i) APPLICANT: Jean, Francois Thomas, Gary
- (ii) TITLE OF INVENTION: Methods and Reagents for Inhibiting Furin Endoprotease
- (iii) NUMBER OF SEQUENCES: 6
- (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: McDonnell Boehnen Hulbert & Berghoff
 - (B) STREET: 300 South Wacker Drive
 - (C) CITY: Chicago (D) STATE: IL

 - (E) COUNTRY: USA
 - (F) ZIP: 60606
- (v) COMPUTER READABLE FORM:

 - (A) MEDIUM TYPE: Floppy disk (B) COMPUTER: IBM PC compatible
 - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.25
- (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER: US (B) FILING DATE: 08-APR-1999

 - (C) CLASSIFICATION:
- (viii) ATTORNEY/AGENT INFORMATION:

 - (A) NAME: Noonan, Kevin E
 (B) REGISTRATION NUMBER: 35,303
 - (C) REFERENCE/DOCKET NUMBER: 92,448-H
 - (ix) TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: 312-913-0001 (B) TELEFAX: 312-913-0002

 - (C) TELEX:
- (2) INFORMATION FOR SEQ ID NO:1:
 - (i) SEQUENCE CHARACTERISTICS:
 (A) LENGTH: 394 amino acids
 (B) TYPE: amino acid

 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: protein
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified site
 - (B) LOCATION: 355..358
 - (C) OTHER INFORMATION: /label=Variant / note="The amino acid sequence is the amino acid sequence of the modified alpha-1-antitrypsin protein, alpha-1-antitrypsin Portland."
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:
 - Glu Asp Pro Gln Gly Asp Ala Ala Gln Lys Thr Asp Thr Ser His His
 - Asp Gln Asp His Pro Thr Phe Asn Lys Ile Thr Pro Asn Leu Ala Glu
 - Phe Ala Phe Ser Leu Tyr Arg Gln Leu Ala His Gln Ser Asn Ser Thr

Asn Ile Phe Phe Ser Pro Val Ser Ile Ala Thr Ala Phe Ala Met Leu Ser Leu Gly Thr Lys Ala Asp Thr His Asp Glu Ile Leu Glu Gly Leu Asn Phe Asn Leu Thr Gln Ile Pro Glu Ala Gln Ile His Glu Gly Phe Gln Glu Leu Leu Arg Thr Leu Asn Gln Pro Asp Ser Gln Leu Gln Leu 105 Thr Thr Gly Asn Gly Leu Phe Leu Ser Gln Gly Leu Lys Leu Val Asp Lys Phe Leu Glu Asp Val Lys Lys Leu Tyr His Ser Glu Ala Phe Thr Val Asn Phe Gly Asp Thr Glu Gln Ala Lys Lys Gln Ile Asn Asp Tyr Val Glu Lys Gly Thr Gln Gly Lys Ile Val Asp Leu Val Lys Glu Leu Asp Arg Asp Thr Val Phe Ala Leu Val Asn Tyr Ile Phe Phe Lys Gly Lys Trp Glu Arg Pro Phe Glu Val Lys Asp Thr Glu Glu Glu Asp Phe His Val Asp Gln Val Thr Thr Val Lys Val Pro Met Met Lys Arg Leu Gly Met Phe Asn Ile Gln His Cys Lys Leu Ser Ser Trp Val Leu Leu Met Lys Tyr Leu Gly Asn Ala Thr Ala Ile Phe Phe Leu Pro Asp 250 Glu Gly Lys Leu Gln His Leu Glu Asn Glu Leu Thr His Asp Ile Ile 260 Thr Lys Phe Leu Glu Asn Glu Asp Arg Arg Ser Ala Ser Leu His Leu Pro Lys Leu Ser Ile Thr Gly Thr Tyr Asp Leu Lys Ser Val Leu Gly 295 Gln Leu Gly Ile Thr Lys Val Phe Ser Asn Gly Ala Asp Leu Ser Gly Val Thr Glu Glu Ala Pro Leu Lys Leu Ser Lys Ala Val His Lys Ala Val Leu Thr Ile Asp Glu Lys Gly Thr Glu Ala Ala Gly Ala Met Phe Leu Glu Arg Ile Pro Arg Ser Ile Pro Pro Glu Val Lys Phe Asn Lys Pro Phe Val Phe Leu Met Ile Glu Gln Asn Thr Lys Ser Pro Leu Phe 375 Met Gly Lys Val Val Asn Pro Thr Gly Lys

- (2) INFORMATION FOR SEQ ID NO:2:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 4 amino acids
 (B) TYPE: amino acid
 (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

Ala Ile Pro Met

- (2) INFORMATION FOR SEQ ID NO:3:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 4 amino acids (B) TYPE: amino acid

 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

Arg Ile Pro Arg

- (2) INFORMATION FOR SEQ ID NO:4:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 4 amino acids
 - (B) TYPE: amino acid (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified site
 (B) LOCATION: 2..3

 - (C) OTHER INFORMATION: /label=Variable site / note="The amino acid Xaa at position 2 can be any amino acid; the amino acid Xaa at position 3 can be any amino acid."
 - (xi) SEQUENCE DESCRIPTION: SEO ID NO:4:

Arg Xaa Xaa Arg

- (2) INFORMATION FOR SEQ ID NO:5:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 4 amino acids (B) TYPE: amino acid

 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:

- (A) NAME/KEY: Modified site
 (B) LOCATION: 2..3
- (C) OTHER INFORMATION: /label=Variable site / note="The amino acid Xaa at position 2 can be any amino acid."
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

Arg Xaa Pro Arg

(2) INFORMATION FOR SEQ ID NO:6:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 394 amino acids
 (B) TYPE: amino acid
 - TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (ix) FEATURE:
 - (A) NAME/KEY: Modified site (B) LOCATION: 355..358

 - (C) OTHER INFORMATION: /label=Variant / note="The amino acid sequence is the amino acid sequence of the modified alpha-1-antitrypsin protein, alpha-1-antitrypsin Pittsburgh.
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:
- Glu Asp Pro Gln Gly Asp Ala Ala Gln Lys Thr Asp Thr Ser His His
- Asp Gln Asp His Pro Thr Phe Asn Lys Ile Thr Pro Asn Leu Ala Glu 20 25 30
- Phe Ala Phe Ser Leu Tyr Arg Gln Leu Ala His Gln Ser Asn Ser Thr
- Asn Ile Phe Phe Ser Pro Val Ser Ile Ala Thr Ala Phe Ala Met Leu
- Ser Leu Gly Thr Lys Ala Asp Thr His Asp Glu Ile Leu Glu Gly Leu
- Asn Phe Asn Leu Thr Gln Ile Pro Glu Ala Gln Ile His Glu Gly Phe
- Gln Glu Leu Leu Arg Thr Leu Asn Gln Pro Asp Ser Gln Leu Gln Leu 105
- Thr Thr Gly Asn Gly Leu Phe Leu Ser Gln Gly Leu Lys Leu Val Asp
- Lys Phe Leu Glu Asp Val Lys Lys Leu Tyr His Ser Glu Ala Phe Thr
- Val Asn Phe Gly Asp Thr Glu Gln Ala Lys Lys Gln Ile Asn Asp Tyr 150
- Val Glu Lys Gly Thr Gln Gly Lys Ile Val Asp Leu Val Lys Glu Leu
- Asp Arg Asp Thr Val Phe Ala Leu Val Asn Tyr Ile Phe Phe Lys Gly

 Lys
 Trp
 Glu 195
 Arg
 Pro
 Phe
 Glu 200
 Lys
 Asp
 Thr
 Glu 205
 Glu Asp
 Phe
 Phe
 200
 Lys
 Asp
 Thr
 Glu Asp
 Phe
 Asp
 Glu 7
 Thr
 Thr
 Lys
 Lys
 Val
 Pro
 Met 220
 Met Lys
 Arg
 Leu 220

 Gly
 Met
 Phe
 Asn
 Ile
 Gln His
 Cys
 Lys
 Lys
 Ser
 Ser
 Trp
 Val
 Leu 240

 Leu
 Met
 Lys
 Lys
 Leu Gly
 Asn
 Ala
 Thr
 Ala
 Ile
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 Leu Pro
 Asp
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 Leu
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 Leu Gly
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